

# **RELATIVE BIOAVAILABILITY OF ARSENIC IN SOILS FROM BUTTE, MONTANA**

Stan W. Casteel, DVM, PhD, DABVT

Principal Investigator

Tim Evans, DVM

Steve Morris, M.S.

Co-Investigators

Veterinary Medical Diagnostic Laboratory

College of Veterinary Medicine

University of Missouri, Columbia

Columbia, Missouri

Susan Griffin, PhD, DABT

Study Design and Technical Advisor

US Environmental Protection Agency – Region 8

Denver, Colorado

William J. Brattin, PhD

Angela M. Wahlquist, MS

Technical Consultants

Syracuse Research Corporation

Denver, Colorado

March 2003

## EXECUTIVE SUMMARY

The gastrointestinal absorption of arsenic from soil samples collected from residential properties at the Butte Arsenic Superfund site was measured using young swine. Test materials include composite soil with arsenic concentrations of 234 ppm (TM1) and 367 ppm (TM2). Groups of animals (four animals per dose group) were given oral doses of reference material (sodium arsenate) or test material twice a day for 12 days. Urine excreted by each animal was collected on days 6-7, 8-9 and 10-11. The urinary excretion fraction (UEF) (the ratio of the amount excreted per 48 hours divided by the dose given per 48 hours) was calculated for each test material using linear regression analysis. The relative bioavailability (RBA) of arsenic in test material compared to that in sodium arsenate (abbreviated NaAs) was calculated as:

$$RBA = \frac{UEF(test\ material)}{UEF(NaAs)}$$

The results are summarized below:

| Test Material | RBA (90% CI) |
|---------------|--------------|
| TM1           | 17% (14-22%) |
| TM2           | 22% (17-29%) |

CI = Confidence Interval

Using sodium arsenate as a relative frame of reference, the RBA estimate for TM1 is 17% and 22% for TM2. These RBA estimates are significantly lower than the default value of 80%-100% that is usually employed for arsenic in soil when reliable site-specific data are lacking. This indicates that the arsenic in these soil samples is not as well absorbed as soluble arsenic. Use of these data is likely to improve the accuracy of risk estimates for humans who may incidentally ingest these soils.

## TABLE OF CONTENTS

|  |    |
|--|----|
| EXECUTIVE SUMMARY .....  | i  |
| 1.0 INTRODUCTION .....   | 1  |
| 2.0 STUDY DESIGN.....  | 3  |
| 2.1 Test Materials.....  | 3  |
| 2.1.1 Test Material Selection and Preparation .....                | 3  |
| 2.1.2 Detailed Characterization of Test Materials .....            | 3  |
| 2.2 Experimental Animals .....                                     | 6  |
| 2.3 Diet.....  | 7  |
| 2.4 Dosing .....   | 7  |
| 2.5 Collection and Preparation of Samples .....                    | 7  |
| 2.6 Arsenic Analysis .....   | 8  |
| 2.6.1 Urine .....  | 8  |
| 2.6.2 Feces .....  | 9  |
| 3.0 DATA ANALYSIS.....   | 10 |
| 4.0 RESULTS .....  | 12 |
| 4.1 Clinical Signs .....   | 12 |
| 4.2 Data Exclusions .....  | 12 |
| 4.3 Urinary Excretion Fractions and Relative Bioavailability ..... | 12 |
| 4.4 Fecal Excretion and Mass Balance .....                         | 13 |
| 5.0 DISCUSSION AND RECOMMENDATIONS.....                            | 14 |
| 6.0 REFERENCES .....   | 15 |

## **LIST OF TABLES**

| <b>TABLE</b> | <b>TITLE</b>                  |
|--------------|-------------------------------|
| 2-1          | Study Design                  |
| 2-2          | Composition of Test Materials |

## **LIST OF FIGURES**

| <b>FIGURE</b> | <b>TITLE</b>  |
|---------------|---|
| 2-1           | Body Weights of Test Animals                            |
| 2-2           | Performance Evaluation Samples                          |
| 2-3           | Blind Duplicate Samples for Urine                       |
| 3-1           | Conceptual Model for Arsenic Toxicokinetics             |
| 4-1           | Urinary Excretion of Arsenic from Sodium Arsenate       |
| 4-2           | Urinary Excretion of Arsenic from Test Material 1       |
| 4-3           | Urinary Excretion of Arsenic from Test Material 2       |
| 4-4           | Urinary Excretion Fraction vs. Fecal Excretion Fraction |

## **LIST OF APPENDICES**

### **APPENDIX A          DETAILED STUDY RESULTS**

|           |   |
|-----------|---|
| Table A-1 | Schedule                                    |
| Table A-2 | Group Assignments                           |
| Table A-3 | Body Weights and Administered Doses, by Day |
| Table A-4 | Body Weight Adjusted Doses                  |
| Table A-5 | Urine Volumes – 48 Hour Collections         |
| Table A-6 | Fecal Weights – 48 Hour Collections         |
| Table A-7 | Urine Analytical Results                    |
| Table A-8 | Feces Analytical Results                    |

### **APPENDIX B          DETAILED ARSENIC SPECIATION RESULTS**

## RELATIVE BIOAVAILABILITY OF ARSENIC IN SOILS FROM BUTTE, MONTANA

### 1.0 INTRODUCTION

Accurate assessment of the health risks resulting from oral exposure to arsenic requires knowledge of the amount of arsenic absorbed from the gastrointestinal tract into the body. This information on absorption may be described either in absolute or relative terms:

Absolute Bioavailability (ABA) is the ratio of the amount of arsenic absorbed to the amount of arsenic ingested:

$$ABA = \frac{\text{Absorbed Dose}}{\text{Ingested Dose}}$$

This ratio is also referred to as the oral absorption fraction ( $AF_o$ ).

Relative Bioavailability (RBA) is the ratio of the absolute bioavailability of arsenic present in some test material to the absolute bioavailability of arsenic in some appropriate reference material:

$$RBA = \frac{ABA (\text{test material})}{ABA (\text{reference material})}$$

Usually the form of arsenic used as the reference material is an arsenic compound dissolved in water or some readily soluble form (e.g., sodium arsenate) that is expected to completely dissolve when ingested.

For example, if 100 µg of arsenic dissolved in drinking water were ingested and a total of 90 µg were absorbed into the body, the ABA would be 0.90 (90%). Likewise, if 100 µg of arsenic contained in soil were ingested and 30 µg were absorbed into the body, the ABA for soil would be 0.30 (30%). If the arsenic dissolved in water were used as the reference substance for describing the relative amount of arsenic absorbed from soil, the RBA would be 0.30/0.90, or 0.33 (33%).

#### Using Relative Bioavailability Data to Improve Risk Calculations for Arsenic

When reliable data are available on the relative bioavailability of arsenic in a site medium (e.g., soil), this information can be used to adjust the default toxicity values ( $RfD_{IRIS}$ ,  $SF_{IRIS}$ ) for arsenic to account for differences in absorption between arsenic ingested in water ( $RBA_w$ ) and arsenic ingested in site media, as follows:

$$RfD_{adj} = \frac{RfD_{IRIS}}{RBA_w}$$

$$SF_{adj} = SF_{IRIS} \cdot RBA_w$$

Alternatively, it is also acceptable to adjust the dose (rather than the toxicity factors) as follows:

$$Dose_{adj} = Dose_{default} \cdot RBA_w$$

This dose adjustment is mathematically equivalent to adjusting the toxicity factors as described above.

### Purpose of this Study

Investigations performed at the Butte Priority Soils Operable Unit in Butte, Montana, have revealed that some residential properties have yard soil that is contaminated with elevated levels of arsenic. This study was performed in order to obtain site-specific data on the relative bioavailability of arsenic in yard soils in order to help improve the accuracy of risk calculations for residents who may be exposed to arsenic in soil.

## 2.0 STUDY DESIGN

This investigation of arsenic relative bioavailability was performed according to the basic design presented in Table 2-1. As shown, the study investigated arsenic absorption from sodium arsenate (the reference material) and from two site-specific soils (the test materials), each administered to groups of animals at three different dose levels for 12 days (a detailed schedule is presented in Appendix A, Table A-1). Additionally, the study included a non-treated group to serve as a control for determining background arsenic levels. All doses were administered orally.

### 2.1 Test Materials

#### 2.1.1 Test Material Description and Preparation

The test materials used in this investigation are two soil samples from Butte, Montana. Test Material 1 (USEPA sample number 8-37926) has been tested previously in the swine bioassay system (USEPA, 1996), and sufficient material existed to repeat the analysis using the same material. This soil sample is a composite collected from the Butte Priority Soils Operable Unit (BPSOU) of the Silver Bow Creek/ Butte Area NPL Site in Butte, Montana. The sampling investigation focused on four source areas: the Little Mina-1, Little Mina-2, West Ruby, and North Emma waste rock dumps. At each source area, five sub-samples were collected and composited, and these were then further composited across source areas to yield the sample used in the study.

Test Material 2 (USEPA sample number BPSOU-0501-ASBIO) was collected by CDM in May 2001 (CDM Federal, 2001). This soil sample is a composite collected from a residential property located adjacent to a railroad grade in Butte, Montana. A total of 5 soil samples from this property were combined in order to prepare the arsenic bioavailability composite sample.

Both composite samples were prepared for administration to the animals by air-drying (maximum temperature = 40°C) followed by sieving through a nylon mesh to yield particles less than about 250 µm. This was done because it is believed that fine particles are most likely to adhere to the hands and be ingested by hand-to-mouth contact, and are most likely to be available for absorption. Grinding was not employed.

#### 2.1.2 Detailed Characterization of Test Materials

##### Arsenic Concentration

Aliquots of each test material were analyzed for arsenic by inductively coupled plasma (ICP) spectroscopy. The results from these analyses are presented below.

| Test Material | Sample ID #                 | Arsenic Concentration (mg/kg) |
|---------------|-----------------------------|-------------------------------|
| 1             | 8-37926 (previously tested) | 234*                          |
| 2             | BPSOU-0501-ASBIO            | 367**                         |

\* Based on quadruplicate analyses via ICP

\*\* Based on triplicate analyses via ICP

### Concentration of Other Inorganics

Each sample was analyzed for EPA's Target Analyte List (TAL) of inorganic chemicals. Results are shown in Table 2-2.

### Particle Speciation, Size, and Matrix Association

Each test material was characterized by electron microprobe analysis (EMPA) in order to identify the different mineral forms of arsenic that were present in each sample and to estimate how much of the total arsenic was present in each form. In addition, the size distribution of the particles was characterized along with the matrix association of each particle. The detailed data are presented in Appendix B and the results are summarized below.

#### *Arsenic Phases*

The following table lists the different arsenic phases observed in the two test materials and gives the relative mass of arsenic (RMA) for each phase in each test material. The RMA is the estimated percentage of the total arsenic in a sample that is present in a particular phase.

| Arsenic Phase                     | Test Material |      |
|-----------------------------------|---------------|------|
|                                   | TM1           | TM2  |
| FeAs Sulfate                      | 53%           | 18%  |
| Clays                             | --            | 0%   |
| FeAs Oxide                        | 20%           | 39%  |
| MnAs Oxide                        | 16%           | --   |
| As Phosphate                      | 8%            | --   |
| AgAsS                             | 2%            | --   |
| Sulfosalts                        | --            | 42%  |
| Pyrite                            | --            | 0.1% |
| AsMSO <sub>4</sub>                | --            | 0.3% |
| Slag                              | --            | 0.0% |
| Barite                            | 0.1%          | --   |
| Total Number of Particles Counted | 636           | 137  |

As seen, arsenic is primarily associated with FeAs sulfate in TM1 and with sulfosalts and FeAs oxide in TM2. These differences in mineral phase may influence the RBA of the arsenic in the materials.

It is important to note that these quantitative estimates of relative arsenic mass are based on examination of a limited number of arsenic-bearing particles in each sample, particularly for Test

Material 2 (N = 137). Consequently, the quantitative values reported should not be considered to be highly precise, and apparent differences between samples may be partly due to random variation in the analysis rather than authentic differences in composition.

### *Particle Size Distribution*

Particle size is a potentially important contributor to RBA because the fraction of a particle that undergoes dissolution in gastrointestinal fluids is likely related to the surface area to volume ratio (this ratio is larger for small particles than large particles). The distribution of particle sizes for arsenic-bearing grains in these test materials is summarized below:

| Test Material | Percent of Particles by Size Class |                      |                    |
|---------------|------------------------------------|----------------------|--------------------|
|               | 0-25 $\mu\text{m}$                 | 26-100 $\mu\text{m}$ | >100 $\mu\text{m}$ |
| TM1           | 49%                                | 40%                  | 11%                |
| TM2           | 57%                                | 39%                  | 4%                 |

As seen, approximately half of the particles in each test material are very small (25  $\mu\text{m}$  or smaller) and the majority (89% to 96%) are 100  $\mu\text{m}$  or less.

### *Matrix Association*

Arsenic-containing particles may be characterized according to their association with other particles into four types, as follows:

| Matrix Association | Description  |
|--------------------|--|
| Liberated          | A grain of arsenic-containing material that is not attached to or contained within any other particle                        |
| Rimming            | Arsenic is present on the outer surface of a particle, usually as a consequence of adsorption or precipitation               |
| Cemented           | The arsenic-containing particle is loosely bound to or associated with other particles or phases that do not contain arsenic |
| Included           | The arsenic-containing particle is entirely contained within another particle  |

In the first three types of matrix association, the arsenic is exposed at the surface of some or all of the particle, and hence the arsenic is available to be dissolved by gastrointestinal fluids. Particles that are fully included in other particles are not exposed to external fluids and are not likely to have high bioavailability. The distribution of matrix associations for arsenic-bearing particles in the test materials from this site is summarized below:

| Test Material | Percent of Particles by Matrix Class |         |          |          |
|---------------|--------------------------------------|---------|----------|----------|
|               | Liberated                            | Rimming | Cemented | Included |
| TM1           | 46%                                  | 6%      | 40%      | 8%       |
| TM2           | 61%                                  | 6%      | 33%      | 0%       |

As seen, relatively few particles are fully included, and 92-100% of the particles are entirely or partially exposed to external fluids. This suggests that the RBA of the arsenic is likely to be determined primarily by mineral phase and/or particle size rather than by matrix association.

### In Vitro Bioaccessibility

The details of the method used to measure the *in vitro* bioaccessibility of arsenic are described in USEPA (1999). In brief, samples of soil are placed in a test fluid designed to be similar to gastric fluid and the fraction of the total amount of arsenic in the sample which dissolves into the fluid under a specified set of conditions (temperature, time, pH) is measured. This fraction of the total arsenic that is solubilized is referred to as the *in vitro* bioaccessibility (IVBA). The IVBA results for the two test materials in this study are summarized below:

| Test Material | IVBA   |
|---------------|--------|
| TM1           | 9% *   |
| TM2           | 13% ** |

\* Based on five analyses

\*\* Based on duplicate analyses

## **2.2 Experimental Animals**

Juvenile swine were selected for use in this study because they are considered to be a good physiological model for gastrointestinal absorption in children (Weis and LaVelle, 1991). The animals were intact males of the Pig Improvement Corporation (PIC) genetically defined Line 26, and were purchased from Chinn Farms, Clarence, MO.

The animals were housed in individual stainless steel cages. All animals were held for several days prior to beginning exposure to test materials to allow them to adapt to their new environment and to ensure that all of the animals were healthy. In order to help minimize weight variations between animals and groups, three animals most different in body weight on day -4 (either heavier or lighter) were also excluded. The remaining animals were assigned to dose groups at random (group assignments are presented in Appendix A, Table A-2). When exposure began (day zero), the animals were about 5-6 weeks old and weighed an average of about 8.6 kg. Animals were weighed every three days during the course of the study. On average, animals gained about 0.3 to 0.4 kg/day, and the rate of weight gain was comparable in all groups. These body weight data are summarized in Figure 2-1 and are also presented in Appendix A, Table A-3.

## **2.3 Diet**

Each day every animal was given an amount of standard swine chow (University Feed Mill S II (2) starter ration without added antibiotics) equal to 5% of the mean body weight of all animals on study. Feed was administered in two equal portions (2.5% of the mean body weight) at 11:00 AM and 5:00 PM daily. Drinking water was provided *ad libitum* via self-activated watering nozzles within each cage.

Based on data from previous arsenic studies, the estimated intake of arsenic in unexposed animals is less than 0.1 µg/kg-day via water and about 10 µg/kg-day via the diet.

## **2.4 Dosing**

Animals were exposed to sodium arsenate (abbreviated in this report as "NaAs") or a test material (site soil) for 12 days, with the dose for each day being administered in two equal portions given at 9:00 AM and 3:00 PM (two hours before feeding). Dose material was placed in the center of a small portion (about 5 grams) of moistened feed (this is referred to as a "doughball"), and this was administered to the animals by hand.

The dose levels administered were based on the arsenic content of the test material, with target doses of 300, 600, and 900 µg/day for NaAs and each test material. The administered arsenic doses are presented in Appendix A, Table A-3, and the body-weight adjusted doses are presented in Appendix A, Table A-4. These actual administered doses were used for all RBA calculations.

## **2.5 Collection and Preparation of Samples**

### Urine

Samples of urine were collected from each animal for three consecutive 48-hour periods, on days 6/7, 8/9, and 10/11, with one exception. It was determined during the first few days of dosing that there were insufficient quantities of Test Material 1 available for dosing according to the protocol. In order to account for this shortage of test material, the dosing for groups 5, 6, and 7 was modified to end one day earlier than originally scheduled. As a result, urine collection for these animals was altered to consist of a 24-hour collection (rather than 48-hour) on day 10, with all other collections being conducted according to schedule.

Urine collections began at 9:00 AM and ended 48 hours later. The urine was collected in a stainless steel pan placed beneath each cage, which drained into a plastic storage bottle. Each collection pan was fitted with a nylon screen to minimize contamination with feces, spilled food, or other debris. Plastic diverters were used to minimize urine dilution with drinking water spilled by the animals from the watering nozzle into the collection pan, although this was not always effective in preventing dilution of the urine with water. Due to the length of the collection period, collection containers were emptied at least twice daily into a separate holding container. This ensured that there was no loss of sample due to overflow.

At the end of each collection period, the urine volume was measured (see Appendix A, A-5) and 60-mL portions were removed for analysis. A separate 250-mL aliquot was retained as an archive sample. Each sample was acidified by the addition of concentrated nitric acid. The samples were stored refrigerated until arsenic analysis.

## Feces

Feces were collected by placing a fine-mesh nylon screen beneath each cage. Samples were transferred from the screen into a storage container twice per day, and the final sample (collected over 48 hours) was weighed (see Appendix A, Table A-6). As for urine, the feces collection for Groups 5, 6, and 7 was altered to consist of a 24-hour collection (rather than 48-hour) on day 10, with all other collections being conducted according to schedule. Aliquots of 20-25 grams of feces were weighed and freeze dried.

## **2.6 Arsenic Analysis**

### **2.6.1 Urine**

Urine samples were arranged in a random sequence and submitted to the laboratory for analysis in a blind fashion.

Details of urine sample preparation and analysis are provided in the study project plan (USEPA, 2001). In brief, 25 mL samples of urine were digested by refluxing and then heating to dryness in the presence of magnesium nitrate and concentrated nitric acid. Following magnesium nitrate digestion, samples were transferred to a muffle furnace and ashed at 500°C. The digested and ashed residue was dissolved in hydrochloric acid and analyzed by the hydride generation technique using a Perkin-Elmer 3100 atomic absorption spectrometer. Preliminary tests of this method established that each of the different forms of arsenic that may occur in urine, including trivalent inorganic arsenic (As+3), pentavalent inorganic arsenic (As+5), mono-methyl arsenic (MMA), and di-methyl arsenic (DMA), are all recovered with high efficiency. Urine analytical results are presented in Appendix A, Table A-7.

## Laboratory Quality Assurance

A number of quality assurance (QA) steps were taken during this project to evaluate the accuracy of the analytical procedures. Steps performed by the analytical laboratory included:

### *Spike Recovery*

Randomly selected urine samples were spiked with known amounts of arsenic (usually 5-10 µg, as sodium arsenate) and the recovery of the added arsenic was measured. Recovery for individual samples typically ranged from 101% to 110%, with an average across all analyses of  $106 \pm 3.2\%$  (N=15).

### *Duplicate Analysis*

The laboratory analyst selected random urine samples for duplicate analysis. Duplicate results typically had a relative percent difference (RPD) of 0-13%, with an average of 2.2% (N=15).

### *Laboratory Control Standards*

Samples of a urine standard were run with each set of test samples. The standard was obtained from ERA (sample number 99106) with a nominal arsenic concentration of 347 µg/L. Results for this standard ranged from 311 to 348 µg/L, with a mean across all samples of  $328 \pm 6.7$  µg/L (N=38).

### *Blanks*

Blank samples run along with each batch of samples never yielded a measurable level of arsenic, with all values being reported as less than 1 µg/L of arsenic.

### Blind Quality Assurance Samples

In addition to these laboratory-sponsored QA procedures, an additional series of QA samples were submitted to the laboratory in a blind fashion. This included a number of Performance Evaluation (PE) samples (control urine spiked with a known amount of arsenic in the form of As<sup>+3</sup>, As<sup>+5</sup>, MMA, or DMA) and a number of blind duplicates.

The results for the PE samples are summarized in Figure 2-2. As seen, good recovery of the arsenic was demonstrated for all standards.

The results for blind duplicates are shown in Figure 2-3. As seen, there was good agreement between results for the duplicate pairs.

Based on the results of all of the quality assurance samples and steps described above, it is concluded that the analytical results for samples of urine are of high quality and are suitable for derivation of reliable estimates of arsenic absorption from test materials.

### **2.6.2 Feces**

After drying, 1.0 gram of fecal material was removed and digested with 10 mL of magnesium nitrate and nitric acid using the same approach as described above for urine. Following digestion, all sample preparation and analytical steps are the same as for urine. Fecal analytical results are presented in Appendix A, Table A-8.

### 3.0 DATA ANALYSIS

Figure 3-1 shows a conceptual model for the toxicokinetic fate of ingested arsenic. Key points of this model are as follow:

- In most animals (including humans), absorbed arsenic is excreted mainly in the urine over the course of several days. Thus, the urinary excretion fraction (UEF), defined as the amount excreted in the urine divided by the amount given, is usually a reasonable approximation of the oral absorption fraction or ABA. However, this ratio will underestimate total absorption, because some absorbed arsenic is excreted in the feces via the bile, and some absorbed arsenic enters tissue compartments (e.g., skin, hair) from which it is cleared very slowly or not at all. Thus the urinary excretion fraction should not be equated with the absolute absorption fraction.
- The relative bioavailability (RBA) of two orally administered materials (e.g., a test material and reference material) can be calculated from the ratio of the urinary excretion fraction of the two materials. This calculation is independent of the extent of tissue binding and of biliary excretion:

$$RBA(test\ vs\ ref) = \frac{AF_o(test)}{AF_o(ref)} = \frac{D \cdot AF_o(test) \cdot K_u}{D \cdot AF_o(ref) \cdot K_u} = \frac{UEF(test)}{UEF(ref)}$$

Based on the conceptual model above, raw data from this study were reduced and analyzed as follows:

- The amount of arsenic excreted in urine by each animal over each collection period was calculated by multiplying the urine volume by the urine concentration:

$$\text{Excreted } (\mu\text{g}/48\text{hr}) = \text{Conc } (\mu\text{g}/\text{L}) \cdot \text{Volume } (\text{L}/48\text{hr})$$

- For each test material, the amount of arsenic excreted by each animal was plotted as a function of the amount administered ( $\mu\text{g}/48$  hours), and the best fit straight line (calculated by linear regression) through the data ( $\mu\text{g}$  excreted per  $\mu\text{g}$  administered) was used as the best estimate of the urinary excretion fraction (UEF).
- The relative bioavailability of arsenic in test material was calculated as:

$$RBA = \text{UEF}(test) / \text{UEF}(\text{NaAs})$$

where sodium arsenate (NaAs) is used as the frame of reference.

- As noted above, each RBA value is calculated as the ratio of two slopes (UEFs), each of which is estimated by linear regression through a set of data points. Because of the variability in the data, there is uncertainty in the estimated slope (UEF) for each material. This uncertainty in the slope is described by the standard error of the mean (SEM) for the slope parameter. Given the best estimate and the SEM for each slope,

the uncertainty in the ratio may be estimated using Monte Carlo simulation. The probability density function (PDF) describing the confidence around each slope term (UEF) was assumed to be characterized by a t-distribution with n-2 degrees of freedom :

$$\frac{UEF(measured) - UEF(true)}{SEM} \sim t_{n-2}$$

For convenience, this PDF is abbreviated T(slope, sem, n), where slope = best estimate of the slope derived by linear regression, sem = standard deviation in the best estimate of the slope, and n = number of data points upon which the regression analysis was performed. Thus, the confidence distribution around each ratio was simulated as:

$$PDF(RBA) = \frac{T(slope, sem, n)_{test}}{T(slope, sem, n)_{ref}}$$

Using this equation, a Monte Carlo simulation was run for each RBA calculation. The 5th and 95th percentile values from the simulated distribution of RBA values were then taken to be the 90% confidence interval for the RBA.

## 4.0 RESULTS

### 4.1 Clinical Signs

The doses of arsenic administered in this study are below a level that is expected to cause toxicological responses in swine, and no clinical signs of arsenic-induced toxicity were noted in any of the animals used in this study.

### 4.2 Data Exclusions

Occasionally, the dilution of urine by spilled water was so large that the concentration of arsenic in the urine could not be quantified. These instances are defined by having a urine arsenic concentration at or below the quantitation limit (2 µg/L) and a total urine volume greater than 5000 mL. When both of these conditions were met, the data were deemed unreliable and excluded from further calculations. In this study, data from two animals in group 1 (pig #157 on days 6/7 and pig# 108 on days 10/11) were deemed unreliable for this reason and excluded. No additional urinary data were excluded.

### 4.3 Urinary Excretion Fractions and Relative Bioavailability

Detailed urinary results from this study are presented in Appendix A. The urinary excretion results for NaAs, Test Material 1, and Test Material 2 are summarized in Figures 4-1, 4-2, and 4-3, respectively. Although there is variability in the data, the dose-response curves are approximately linear, with the slope of the best-fit straight line being equal to the best estimate of the urinary excretion fraction (UEF). This finding is consistent with results from both animals and humans, which suggest that there is no threshold for arsenic absorption or excretion up to doses of at least 5,000 µg/day (USEPA, 1995).

As discussed above, the relative bioavailability of arsenic in a specific test material is calculated as follows:

$$\text{RBA}(\text{test vs NaAs}) = \text{UEF}(\text{test}) / \text{UEF}(\text{NaAs, oral})$$

The following table summarizes the best fit slopes (urinary excretion fractions) for sodium arsenate and each of the test materials, as well as the RBA estimates:

| Test Material          | Slope (UEF) ± SEM  | RBA (90% CI)     |
|------------------------|--------------------|------------------|
| NaAs                   | 0.907 ± 0.110 (43) | [1.00]           |
| TM1 (8-37926)          | 0.154 ± 0.009 (41) | 0.17 (0.14-0.22) |
| TM2 (BPSOU-0501-ASBIO) | 0.202 ± 0.018 (43) | 0.22 (0.17-0.29) |

CI = Confidence Interval

As seen, using sodium arsenate as a relative frame of reference, the RBA estimate is 17% for TM1 and 22% for TM2. These RBA estimates are significantly lower than the default value of 80%-100% that is usually employed for arsenic in soil when reliable site-specific data are

lacking. This indicates that the arsenic in these soil samples is not as well absorbed as soluble arsenic, and it is appropriate to take this into account when evaluating potential risks to humans from incidental ingestion of these soils.

#### 4.4 Fecal Excretion and Mass Balance

As shown in Figure 3-1, the amount of arsenic excreted in the feces is the sum of that which is ingested but never absorbed and that which is absorbed and then secreted in bile back into the intestines. Assuming that biliary excretion of absorbed arsenic is a relatively minor metabolic pathway, then the amount of arsenic excreted in the feces is expected to be high when the urinary excretion fraction (and hence the RBA) is low.

Detailed fecal data from this study are presented in Appendix A, Table A-8. Figure 4-4 shows the fecal excretion fraction (defined as the mass of arsenic excreted in feces in 48 hours divided by the oral dose of arsenic administered in 48 hours) as a function of the urinary excretion fraction. As seen, there is a clear negative trend, with low urinary excretion being associated with high fecal excretion.

The sum of the two excretion fractions is equal to the total fraction of the administered dose recovered in urine plus feces. These data are summarized below:

| Test Material | UEF   | FEF   | Total |
|---------------|-------|-------|-------|
| NaAs          | 0.907 | 0.051 | 0.958 |
| TM1           | 0.154 | 0.638 | 0.793 |
| TM2           | 0.202 | 0.692 | 0.894 |

As seen, the total fraction of the administered arsenic that was recovered in urine and feces averaged about 88% (range = 79%-96%). This recovery is consistent with most other studies of arsenic excretion in animals (USEPA, 1995).

## **5.0 DISCUSSION AND RECOMMENDATIONS**

The RBA estimates for site soils collected from the Butte study area are about 0.17 and 0.22, with a mean of about 0.20. These values are both substantially less than the default value of 0.8 recommended by USEPA Region 8, supporting the conclusion that arsenic in Butte site soils is not as well absorbed as soluble arsenic. The detailed chemical mechanism accounting for this reduced bioavailability of arsenic in site soils is not known, but almost certainly is related to the chemical form of arsenic in the soils.

As mentioned in Section 2.1.1, Test Material 1 was tested previously in the swine bioassay system (USEPA, 1996) using an older analytical protocol that tended to have low recovery of organic arsenic. During that study, 24-hour urine samples were collected on Days 7 and 14, and the RBA was determined to be 0.06. Although this previous estimate is slightly lower than the estimate obtained in this study using the newer arsenic analysis method ( $\text{MgNO}_3$  digestion) (0.17), both values support the conclusion that the RBA of TM1 is quite low compared to the USEPA default value (0.8). A comparison of the methods and results between the first and second analysis of TM1 is described in greater detail in a separate report (USEPA, 2003).

## 6.0 REFERENCES

CDM Federal. 2001. Data summary report for arsenic bioavailability study soil sampling conducted May 11 and May 17, 2001 in Butte and Walkerville, Montana. Report addressed to Sara Sparks, USEPA Region VIII, by CDM Federal Programs Corporation, October 19, 2001.

USEPA. 2003. Comparison of relative bioavailability of arsenic in soil estimated using data derived with two alternative analytical methods. Report prepared by Syracuse Research Corporation for USEPA Region 8. April 2003 (draft).

USEPA. 2001. Quality Assurance Project Plan. Bioavailability of Arsenic in Soil from Butte, Montana Using Juvenile Swine as an Animal Model. Report prepared by Syracuse Research Corporation for USEPA Region 8. June 2001.

USEPA. 1999. In Vitro Bioaccessibility (ISSI SOP No. ISSI-VBI70-10). USEPA Region VIII Arsenic Bioavailability Study SOP #16. SOP prepared by ISSI Consulting Group, Inc., for USEPA Region VIII, September 1999 (Rev. #0). In: Quality assurance project plan for bioavailability of arsenic in soil from Butte, Montana evaluated using juvenile swine as an animal model; prepared by USEPA Region VIII and Syracuse Research Corporation, June 2001.

USEPA. 1996. Bioavailability of Arsenic in Soil from the Silver Bow Creek/Butte Area NPL Site. Butte, Montana. Report prepared by Roy F. Weston for USEPA Region 8. May 1996.

USEPA. 1995. Bioavailability of Metals in Soils and Solid Wastes. Report prepared by Roy F. Weston, Inc., for USEPA Region 8. February 1995. Document Control Number 4800-045-008.

Weis, C.P., and LaVelle, J.M. 1991. Characteristics to consider when choosing an animal model for the study of lead bioavailability. In: The Proceedings of the International Symposium on the Bioavailability and Dietary Uptake of Lead. Science and Technology Letters 3:113-119.

## TABLES

**TABLE 2-1 STUDY DESIGN**

| Group | Number of Animals | Material Administered | Target Arsenic Dose (ug/kg-day) |
|-------|-------------------|-----------------------|---------------------------------|
| 1     | 3                 | Control               | 0                               |
| 2     | 4                 | NaAs                  | 25                              |
| 3     | 4                 | NaAs                  | 50                              |
| 4     | 4                 | NaAs                  | 75                              |
| 5     | 4                 | Test Material 1       | 25                              |
| 6     | 4                 | Test Material 1       | 50                              |
| 7     | 4                 | Test Material 1       | 75                              |
| 8     | 4                 | Test Material 2       | 25                              |
| 9     | 4                 | Test Material 2       | 50                              |
| 10    | 4                 | Test Material 2       | 75                              |

**TABLE 2-2 COMPOSITION OF TEST MATERIALS**

| Analyte   | Concentration (mg/kg) |       |
|-----------|-----------------------|-------|
|           | TM1                   | TM2   |
| Aluminum  | 7970                  | 14067 |
| Antimony  | 6.2                   | 3.4   |
| Arsenic   | 251*                  | 367** |
| Barium    | 142                   | 211   |
| Beryllium | 0.61                  | 0.78  |
| Cadmium   | 43.1                  | 7.7   |
| Calcium   | 16100                 | 3363  |
| Chromium  | 7.4                   | 25.6  |
| Cobalt    | 9.6                   | 8.7   |
| Copper    | 871                   | 3130  |
| Iron      | 52100                 | 39800 |
| Lead      | 8640                  | 492   |
| Magnesium | 3090                  | 3950  |
| Manganese | 13500                 | 732   |
| Mercury   | 2.1                   | 0.4   |
| Nickel    | 9                     | 12    |
| Potassium | 3640                  | 3680  |
| Selenium  | 0.28                  | 0.91  |
| Silver    | 42.3                  | 8.1   |
| Sodium    | 537                   | 777   |
| Thallium  | 1.8                   | 0.8   |
| Vanadium  | 30                    | 49    |
| Zinc      | 12500                 | 2457  |

\* Based on quadruplicate analyses via ICP

\*\* Based on triplicate analyses via ICP

## FIGURES

**FIGURE 2-1 BODY WEIGHTS OF TEST ANIMALS**

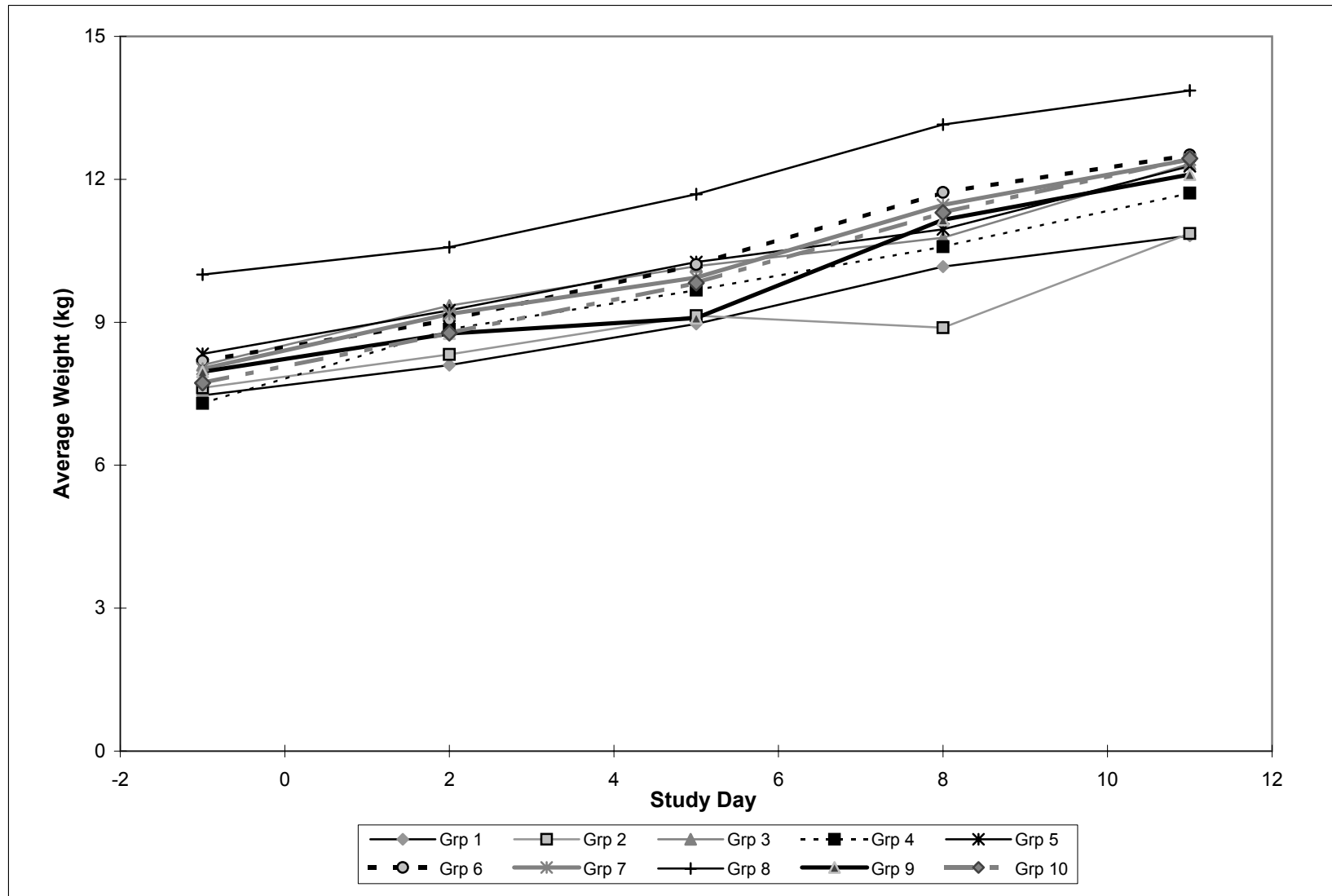
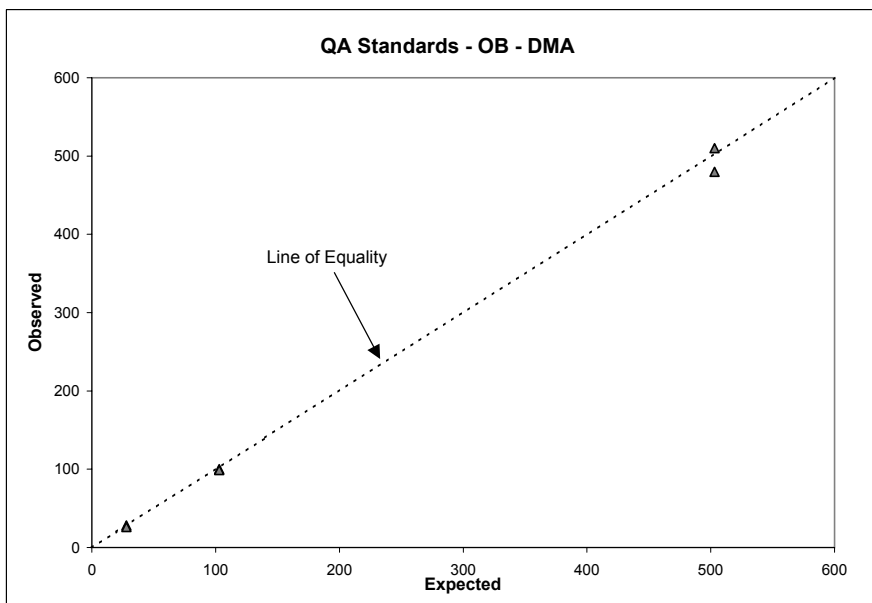
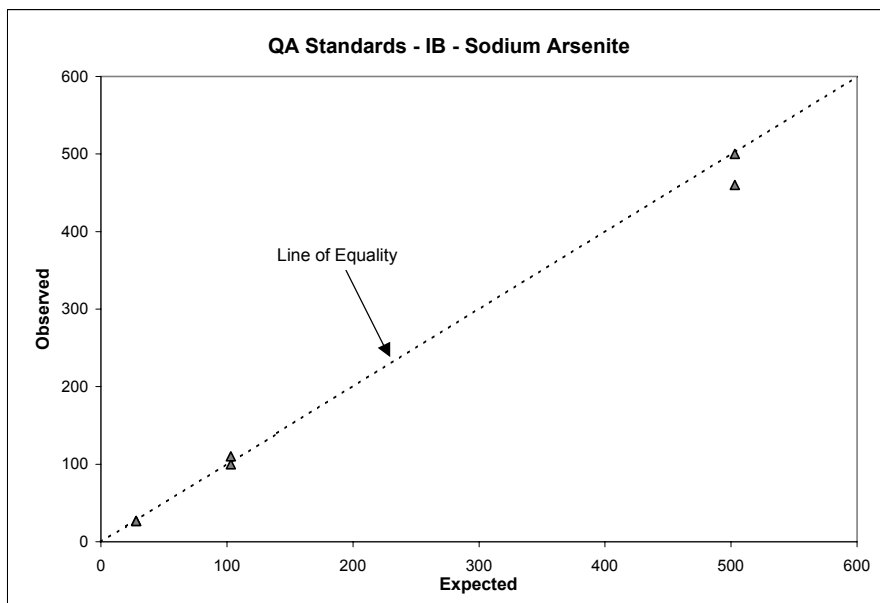
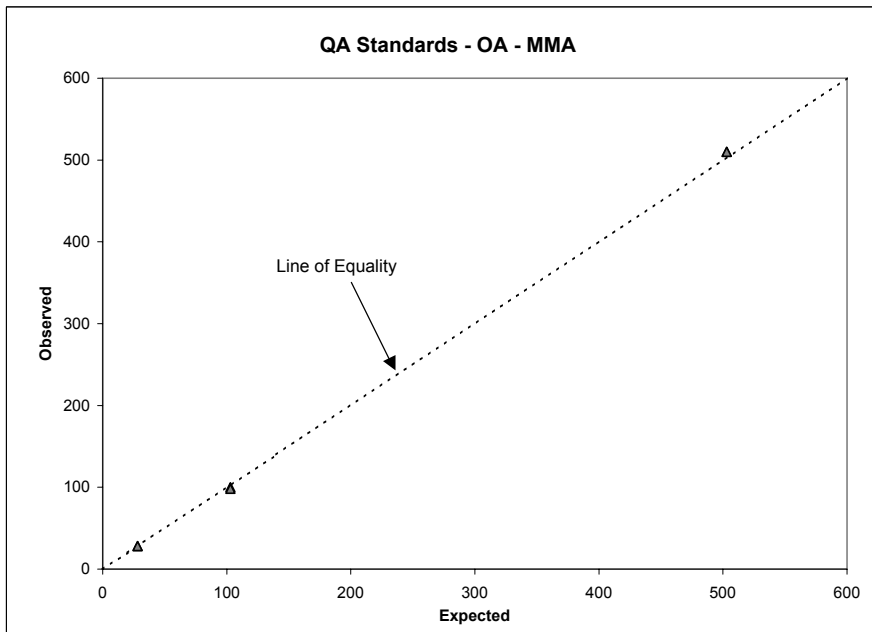
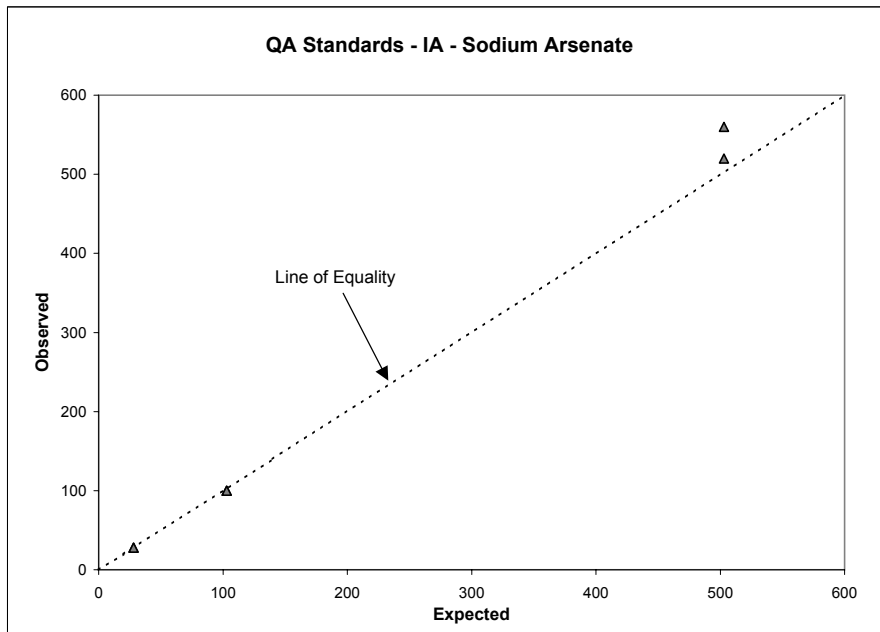
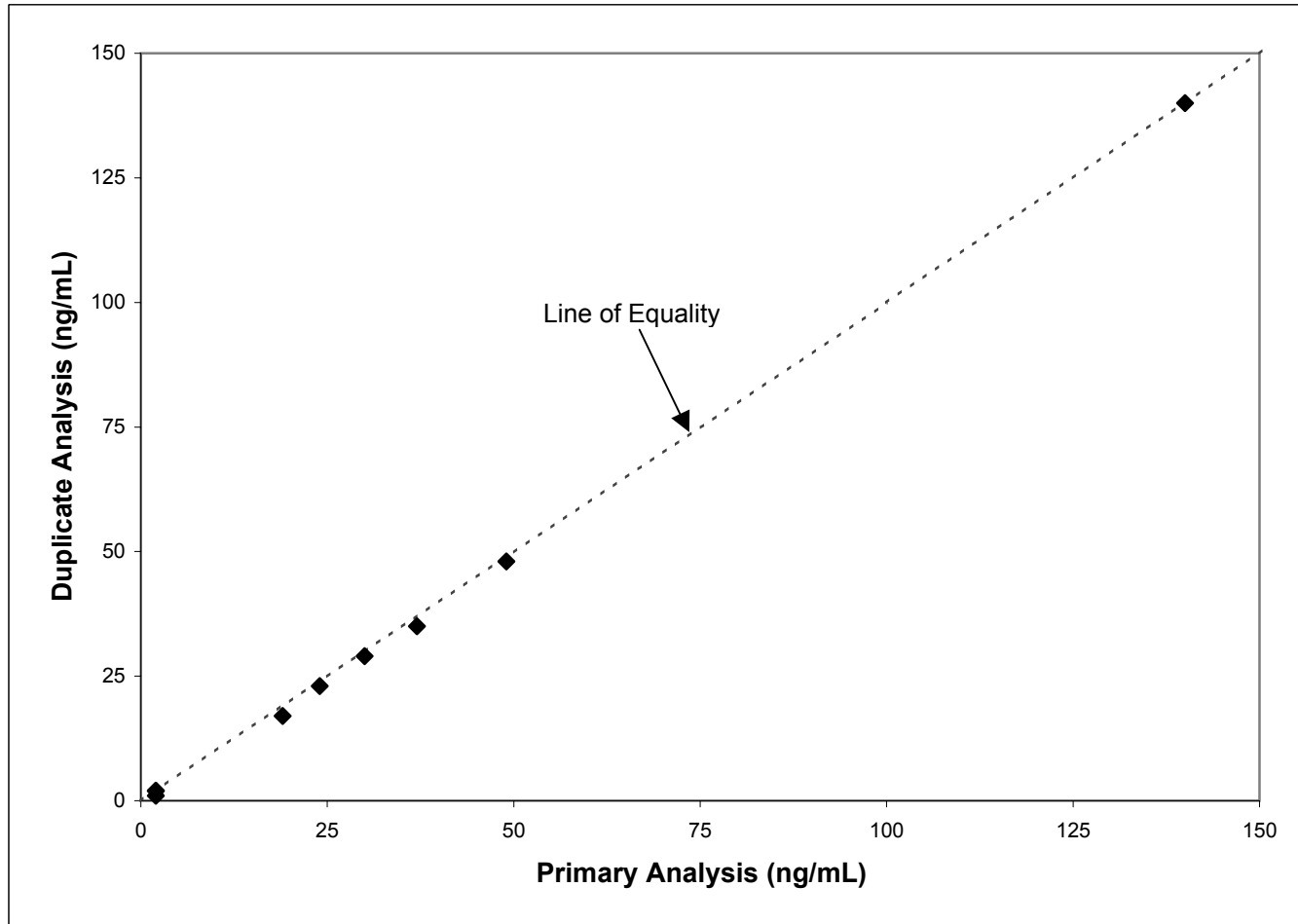


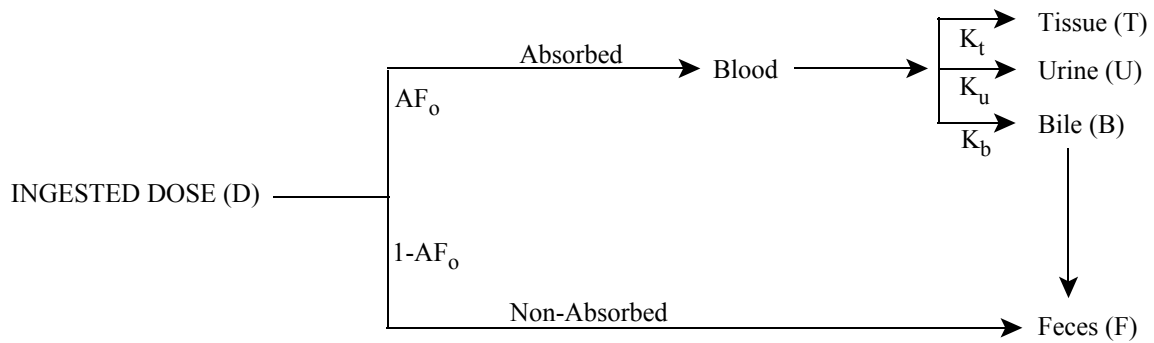
FIGURE 2-2 PERFORMANCE EVALUATION SAMPLES



**FIGURE 2-3 BLIND DUPLICATE SAMPLES FOR URINE**



**Figure 3-1. Conceptual Model for Arsenic Toxicokinetics**



where:

$D$  = Ingested dose (ug)

$AF_o$  = Oral Absorption Fraction

$K_t$  = Fraction of absorbed arsenic which is retained in tissues

$K_u$  = Fraction of absorbed arsenic which is excreted in urine

$K_b$  = Fraction of absorbed arsenic which is excreted in the bile

#### BASIC EQUATIONS:

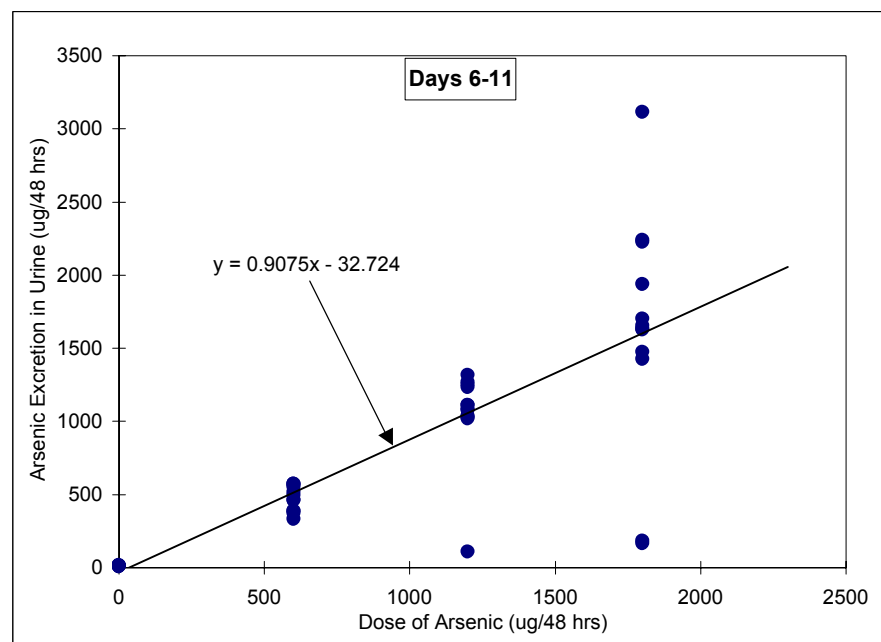
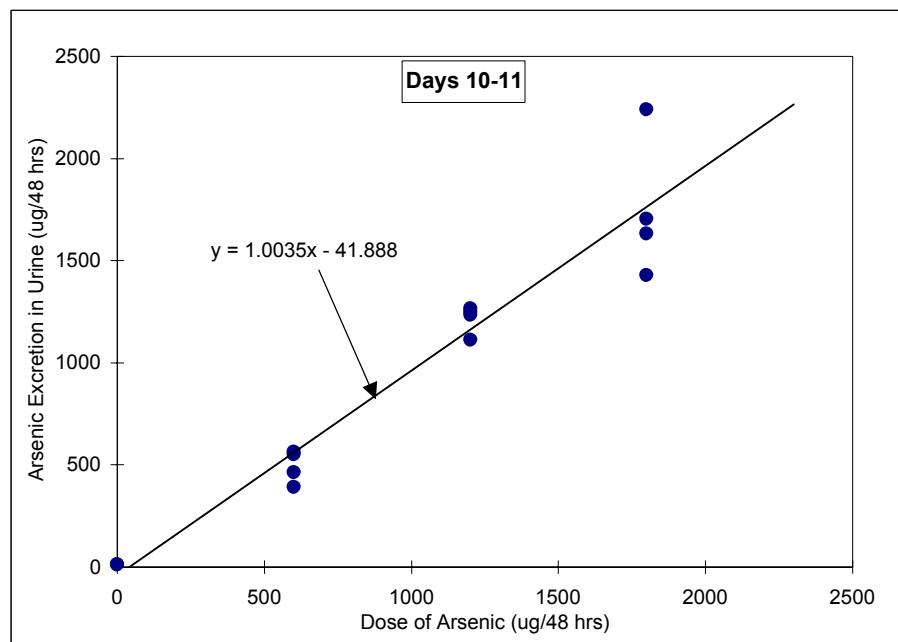
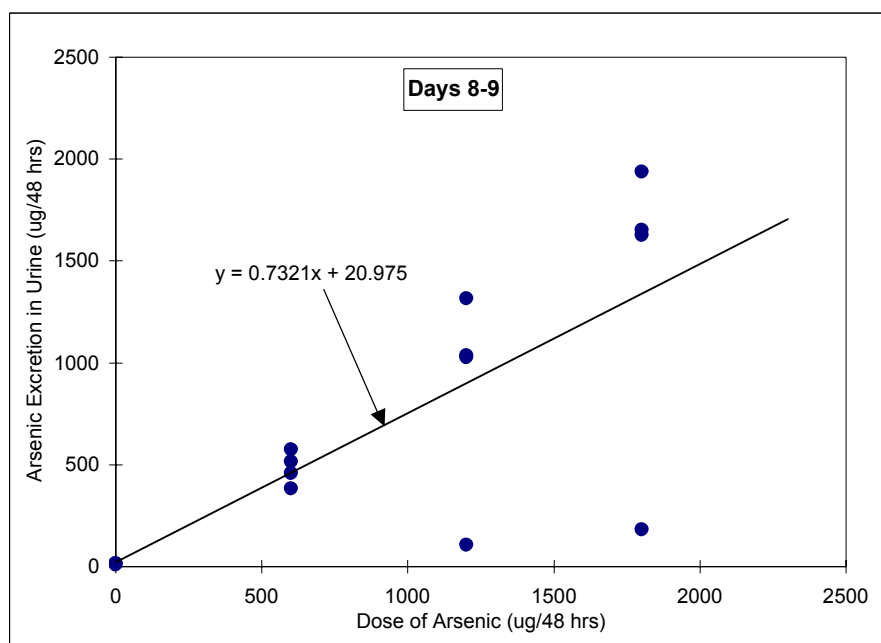
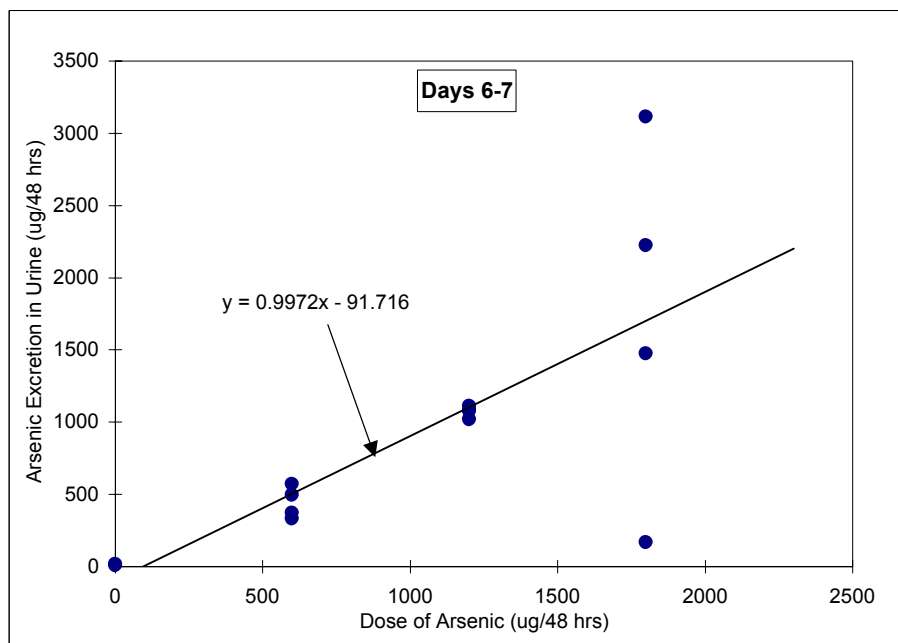
$$\text{Amount Absorbed (ug)} = D \cdot AF_o$$

$$\begin{aligned} \text{Amount Excreted (ug)} &= \text{Amount absorbed} \cdot K_u \\ &= D \cdot AF_o \cdot K_u \end{aligned}$$

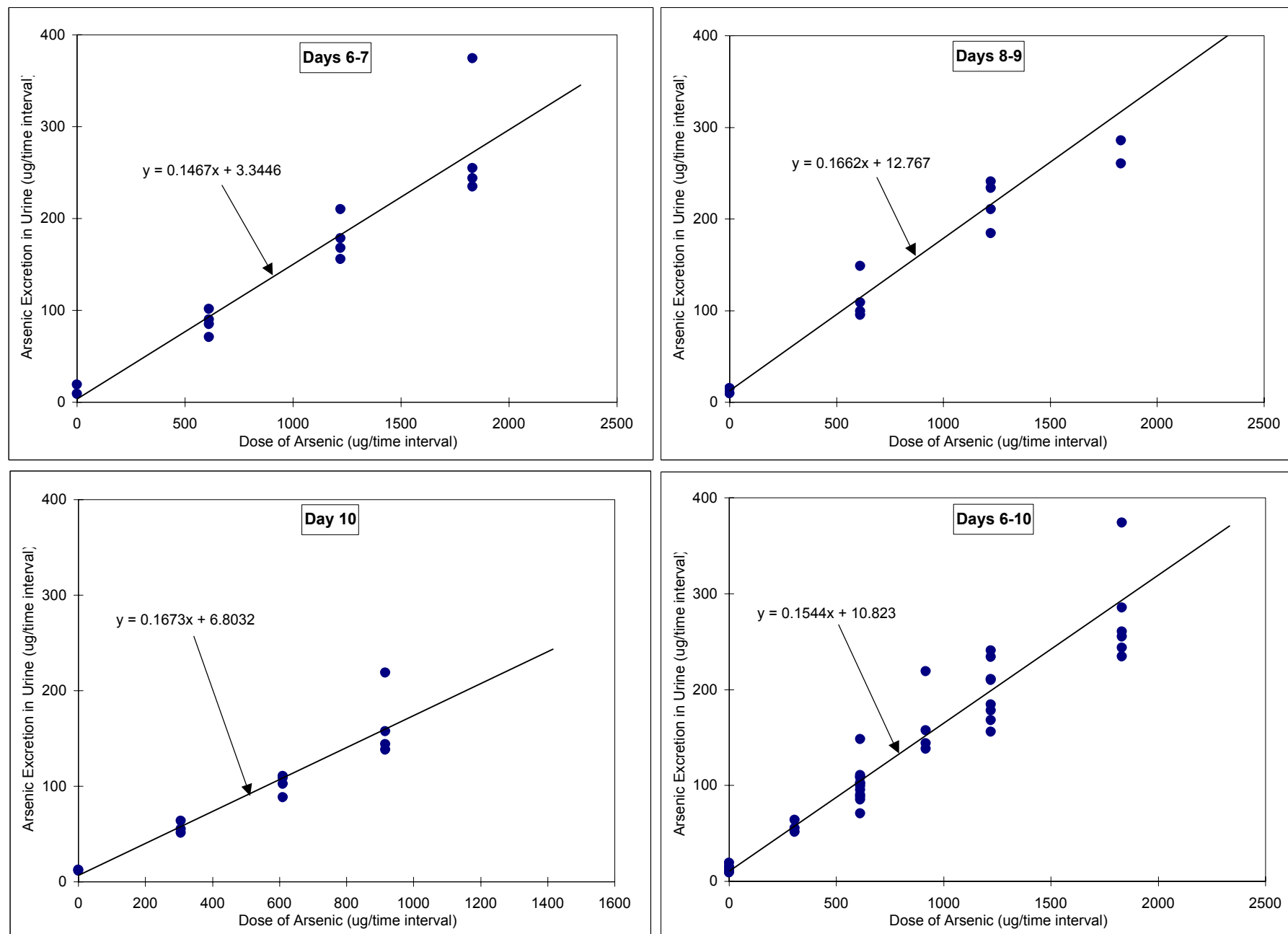
$$\begin{aligned} \text{Urinary Excretion Fraction (UEF)} &= \text{Amount excreted} / \text{Amount Ingested} \\ &= (D \cdot AF_o \cdot K_u) / D \\ &= AF_o \cdot K_u \end{aligned}$$

$$\begin{aligned} \text{Relative Bioavailability (x vs. y)} &= \text{UEF(x)} / \text{UEF(y)} \\ &= (AF_o(x) \cdot K_u) / (AF_o(y) \cdot K_u) \\ &= AF_o(x) / AF_o(y) \end{aligned}$$

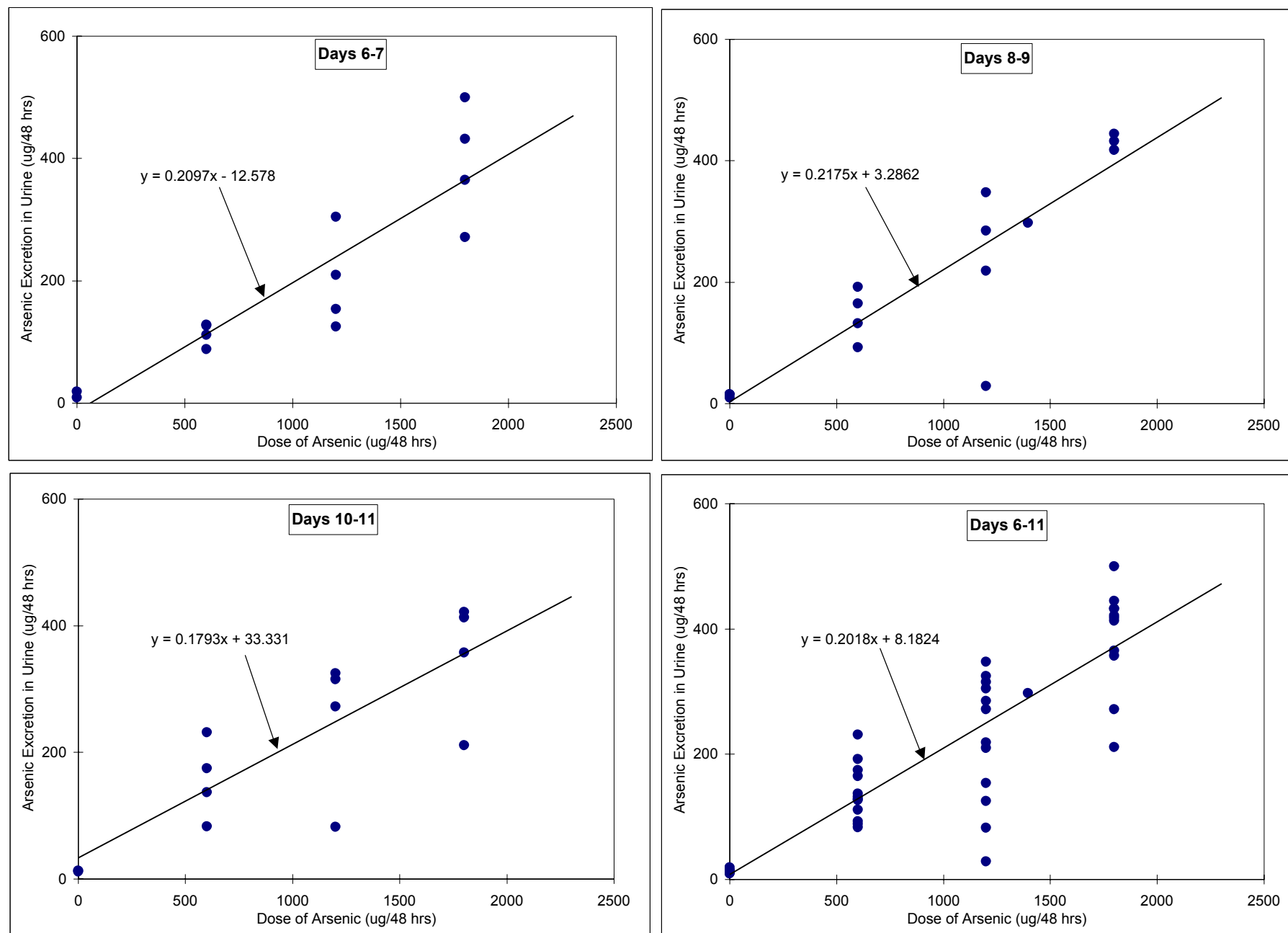
**FIGURE 4-1 URINARY EXCRETION OF ARSENIC FROM SODIUM ARSENATE**



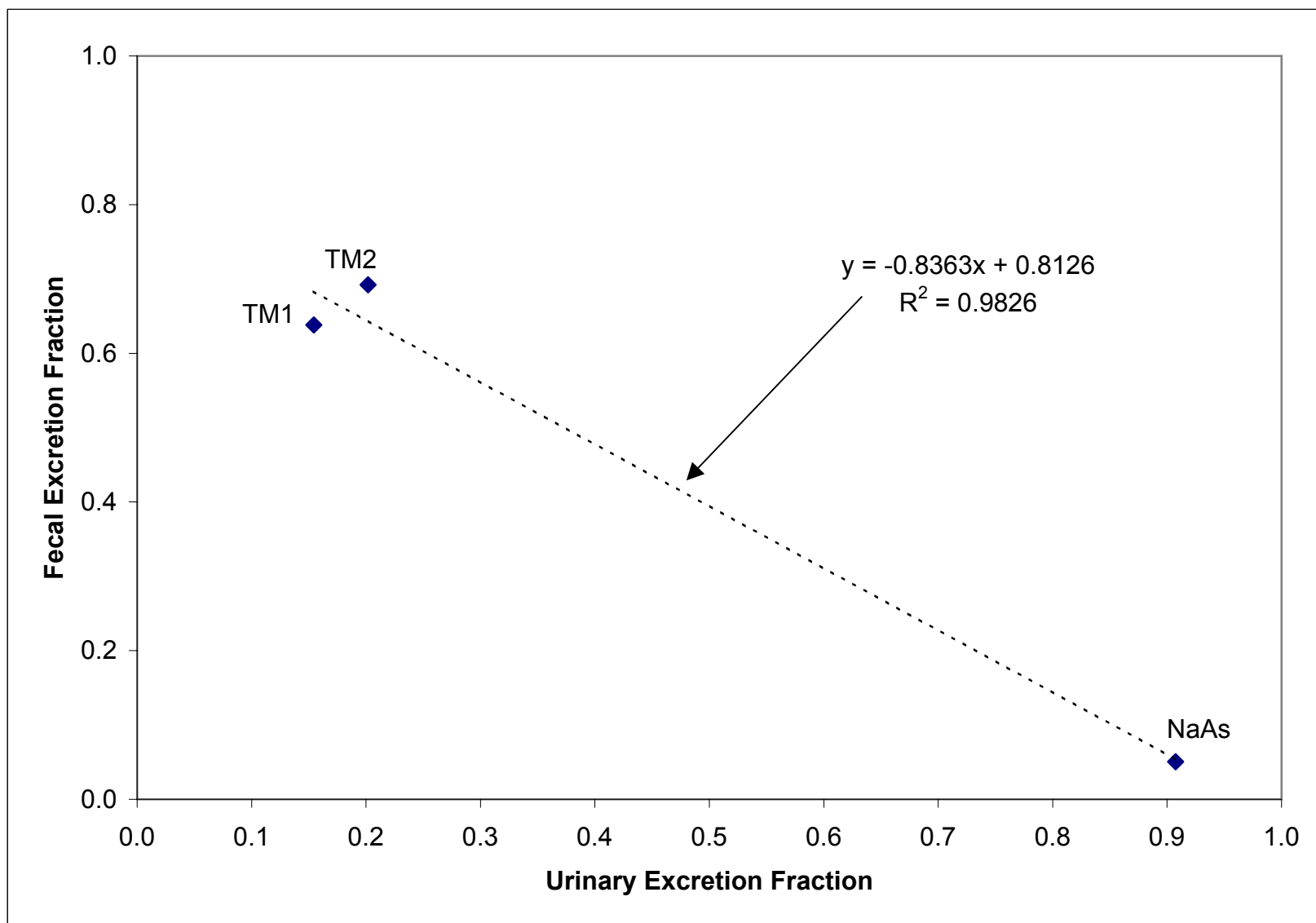
**FIGURE 4-2 URINARY EXCRETION OF ARSENIC FROM TEST MATERIAL 1**



**FIGURE 4-3 URINARY EXCRETION OF ARSENIC FROM TEST MATERIAL 2**



**FIGURE 4-4 URINARY EXCRETION FRACTION VS. FECAL EXCRETION FRACTION**



## **APPENDIX A**

### **DETAILED STUDY RESULTS**

**TABLE A-1 SCHEDULE**

| Study Day | Day       | Date      | Dose Administration | Feed | Weigh | Dose Prep | Cull Pigs/Assign Dose Group | 48 hr Urine Collection | Sacrifice |
|-----------|-----------|-----------|---------------------|------|-------|-----------|-----------------------------|------------------------|-----------|
| -4        | Saturday  | 6/16/2001 |                     | X    | X     |           |                             |                        |           |
| -3        | Sunday    | 6/17/2001 |                     | X    |       |           |                             |                        |           |
| -2        | Monday    | 6/18/2001 |                     | X    |       |           | X                           |                        |           |
| -1        | Tuesday   | 6/19/2001 |                     | X    | X     | X         |                             |                        |           |
| 0         | Wednesday | 6/20/2001 | X                   | X    |       |           |                             |                        |           |
| 1         | Thursday  | 6/21/2001 | X                   | X    |       |           |                             |                        |           |
| 2         | Friday    | 6/22/2001 | X                   | X    | X     |           |                             |                        |           |
| 3         | Saturday  | 6/23/2001 | X                   | X    |       |           |                             |                        |           |
| 4         | Sunday    | 6/24/2001 | X                   | X    |       |           |                             |                        |           |
| 5         | Monday    | 6/25/2001 | X                   | X    | X     |           |                             |                        |           |
| 6         | Tuesday   | 6/26/2001 | X                   | X    |       |           |                             | ↑                      |           |
| 7         | Wednesday | 6/27/2001 | X                   | X    |       |           |                             | ↓                      |           |
| 8         | Thursday  | 6/28/2001 | X                   | X    | X     |           |                             | ↑                      |           |
| 9         | Friday    | 6/29/2001 | X                   | X    |       |           |                             | ↓                      |           |
| 10        | Saturday  | 6/30/2001 | X                   | X    |       |           |                             | ↑                      |           |
| 11        | Sunday    | 7/1/2001  | X                   | X    | X     |           |                             | ↓                      |           |
| 12        | Monday    | 7/2/2001  |                     |      |       |           |                             |                        | X         |

**TABLE A-2 GROUP ASSIGNMENTS**

| Pig Number | Group | Material Administered | Target Dose of Arsenic (ug/kg-day) |
|------------|-------|-----------------------|------------------------------------|
| 108        | 1     | Control               | 0                                  |
| 145        |       |                       |                                    |
| 157        |       |                       |                                    |
| 122        | 2     | NaAs                  | 25                                 |
| 123        |       |                       |                                    |
| 147        |       |                       |                                    |
| 156        |       |                       |                                    |
| 101        | 3     | NaAs                  | 50                                 |
| 115        |       |                       |                                    |
| 119        |       |                       |                                    |
| 151        |       |                       |                                    |
| 121        | 4     | NaAs                  | 75                                 |
| 136        |       |                       |                                    |
| 140        |       |                       |                                    |
| 148        |       |                       |                                    |
| 104        | 5     | TM1                   | 25                                 |
| 106        |       |                       |                                    |
| 128        |       |                       |                                    |
| 155        |       |                       |                                    |
| 103        | 6     | TM1                   | 50                                 |
| 110        |       |                       |                                    |
| 116        |       |                       |                                    |
| 142        |       |                       |                                    |
| 120        | 7     | TM1                   | 75                                 |
| 125        |       |                       |                                    |
| 138        |       |                       |                                    |
| 150        |       |                       |                                    |
| 102        | 8     | TM2                   | 25                                 |
| 114        |       |                       |                                    |
| 117        |       |                       |                                    |
| 126        |       |                       |                                    |
| 112        | 9     | TM2                   | 50                                 |
| 113        |       |                       |                                    |
| 135        |       |                       |                                    |
| 154        |       |                       |                                    |
| 124        | 10    | TM2                   | 75                                 |
| 133        |       |                       |                                    |
| 158        |       |                       |                                    |
| 160        |       |                       |                                    |

**TABLE A-3. BODY WEIGHTS AND ADMINISTERED DOSES, BY DAY**

Body weights were measured on days -1, 2, 5, 8, 11. Weights for other days are estimated, based on linear interpolation between measured values.

| Group | ID # | Day -1  |               | Day 0   |               | Day 1   |               | Day 2   |               | Day 3   |               | Day 4   |               | Day 5   |               | Day 6   |               | Day 7   |               | Day 8   |               | Day 9   |               | Day 10  |               | Day 11  |               |
|-------|------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|---------|---------------|
|       |      | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day | BW (kg) | ug As per day |
| 1     | 108  | 7.35    | 0             | 7.5     | 0             | 7.6     | 0             | 7.75    | 0             | 7.9     | 0             | 8.1     | 0             | 8.3     | 0             | 8.8     | 0             | 9.2     | 0             | 9.65    | 0             | 9.8     | 0             | 10.0    | 0             | 10.15   | 0             |
| 1     | 145  | 7.8     | 0             | 7.9     | 0             | 8.1     | 0             | 8.2     | 0             | 8.6     | 0             | 9.0     | 0             | 9.45    | 0             | 9.8     | 0             | 10.1    | 0             | 10.35   | 0             | 10.7    | 0             | 11.0    | 0             | 11.25   | 0             |
| 1     | 157  | 7.25    | 0             | 7.6     | 0             | 8.0     | 0             | 8.35    | 0             | 8.6     | 0             | 8.9     | 0             | 9.15    | 0             | 9.6     | 0             | 10.1    | 0             | 10.5    | 0             | 10.7    | 0             | 10.9    | 0             | 11.05   | 0             |
| 2     | 122  | 8.1     | 0             | 8.4     | 300           | 8.7     | 300           | 8.95    | 300           | 9.3     | 300           | 9.7     | 300           | 10.05   | 300           | 9.4     | 300           | 8.8     | 300           | 8.15    | 300           | 9.6     | 300           | 11.1    | 300           | 12.5    | 300           |
| 2     | 123  | 7.25    | 0             | 7.6     | 300           | 7.9     | 300           | 8.25    | 300           | 8.6     | 300           | 9.0     | 300           | 9.4     | 300           | 9.7     | 300           | 10.0    | 300           | 10.35   | 300           | 10.7    | 300           | 11.0    | 300           | 11.3    | 300           |
| 2     | 147  | 7.85    | 0             | 8.0     | 300           | 8.1     | 300           | 8.15    | 300           | 8.4     | 300           | 8.6     | 300           | 8.8     | 300           | 8.8     | 300           | 8.8     | 300           | 8.85    | 300           | 9.3     | 300           | 9.8     | 300           | 10.2    | 300           |
| 2     | 156  | 7.3     | 0             | 7.5     | 300           | 7.7     | 300           | 7.95    | 300           | 8.1     | 300           | 8.2     | 300           | 8.3     | 300           | 8.3     | 300           | 8.2     | 300           | 8.2     | 300           | 8.6     | 300           | 9.0     | 300           | 9.45    | 300           |
| 3     | 101  | 7.75    | 0             | 8.4     | 600           | 9.1     | 600           | 9.8     | 600           | 10.2    | 600           | 10.6    | 600           | 10.95   | 600           | 11.2    | 600           | 11.4    | 600           | 11.55   | 600           | 12.2    | 600           | 12.8    | 600           | 13.35   | 600           |
| 3     | 115  | 7.55    | 0             | 8.0     | 600           | 8.4     | 600           | 8.85    | 600           | 9.2     | 600           | 9.5     | 600           | 9.8     | 600           | 9.9     | 600           | 9.9     | 600           | 10      | 600           | 10.6    | 600           | 11.2    | 600           | 11.75   | 600           |
| 3     | 119  | 9       | 0             | 9.3     | 600           | 9.5     | 600           | 9.8     | 600           | 10.0    | 600           | 10.3    | 600           | 10.5    | 600           | 10.6    | 600           | 10.6    | 600           | 10.7    | 600           | 11.5    | 600           | 12.2    | 600           | 12.95   | 600           |
| 3     | 151  | 8.1     | 0             | 8.4     | 600           | 8.7     | 600           | 8.95    | 600           | 9.1     | 600           | 9.3     | 600           | 9.45    | 600           | 9.9     | 600           | 10.4    | 600           | 10.85   | 600           | 11.0    | 600           | 11.1    | 600           | 11.25   | 600           |
| 4     | 121  | 5.15    | 0             | 6.03    | 900           | 6.9     | 900           | 7.8     | 900           | 8.0     | 900           | 8.1     | 900           | 8.3     | 900           | 9.0     | 900           | 9.6     | 900           | 10.3    | 900           | 10.5    | 900           | 10.7    | 900           | 10.85   | 900           |
| 4     | 136  | 9.35    | 0             | 9.5     | 900           | 9.6     | 900           | 9.75    | 900           | 10.2    | 900           | 10.6    | 900           | 10.95   | 900           | 11.2    | 900           | 11.4    | 900           | 11.6    | 900           | 12.0    | 900           | 12.4    | 900           | 12.8    | 900           |
| 4     | 140  | 6.95    | 0             | 7.6     | 900           | 8.2     | 900           | 8.75    | 900           | 9.0     | 900           | 9.2     | 900           | 9.4     | 900           | 9.4     | 900           | 9.4     | 900           | 9.4     | 900           | 10.0    | 900           | 10.6    | 900           | 11.2    | 900           |
| 4     | 148  | 7.75    | 0             | 8.2     | 900           | 8.7     | 900           | 9.1     | 900           | 9.4     | 900           | 9.7     | 900           | 10.05   | 900           | 10.4    | 900           | 10.7    | 900           | 11.05   | 900           | 11.4    | 900           | 11.7    | 900           | 12      | 900           |
| 5     | 104  | 8.55    | 0             | 8.8     | 305           | 9.0     | 305           | 9.15    | 305           | 9.5     | 306           | 9.9     | 306           | 10.3    | 306           | 10.3    | 306           | 10.3    | 306           | 10.3    | 306           | 10.9    | 306           | 11.5    | 306           | 12.15   | 306           |
| 5     | 106  | 8.6     | 0             | 8.7     | 305           | 8.7     | 305           | 8.8     | 305           | 9.2     | 306           | 9.5     | 306           | 9.85    | 306           | 10.5    | 306           | 11.1    | 306           | 11.65   | 306           | 11.9    | 306           | 12.2    | 306           | 12.4    | 306           |
| 5     | 128  | 8.85    | 0             | 9.4     | 305           | 9.9     | 305           | 10.45   | 305           | 10.8    | 306           | 11.1    | 306           | 11.35   | 306           | 11.0    | 306           | 10.7    | 306           | 10.3    | 306           | 11.1    | 306           | 11.9    | 306           | 12.7    | 306           |
| 5     | 155  | 7.35    | 0             | 7.8     | 305           | 8.2     | 305           | 8.6     | 305           | 8.9     | 306           | 9.2     | 306           | 9.55    | 306           | 10.2    | 306           | 10.9    | 306           | 11.55   | 306           | 11.7    | 306           | 11.8    | 306           | 11.85   | 306           |
| 6     | 103  | 7.35    | 0             | 7.6     | 610           | 7.8     | 610           | 8.05    | 610           | 8.4     | 611           | 8.8     | 611           | 9.2     | 611           | 9.7     | 611           | 10.1    | 611           | 10.6    | 611           | 10.9    | 611           | 11.2    | 611           | 11.5    | 611           |
| 6     | 110  | 8.5     | 0             | 9.1     | 610           | 9.7     | 610           | 10.35   | 610           | 10.7    | 611           | 11.1    | 611           | 11.45   | 611           | 12.0    | 611           | 12.5    | 611           | 12.95   | 611           | 13.3    | 611           | 13.6    | 611           | 13.95   | 611           |
| 6     | 116  | 9.6     | 0             | 9.8     | 610           | 10.0    | 610           | 10.15   | 610           | 10.7    | 611           | 11.2    | 611           | 11.75   | 611           | 12.4    | 611           | 13.0    | 611           | 13.6    | 611           | 13.8    | 611           | 14.1    | 611           | 14.3    | 611           |
| 6     | 142  | 7.3     | 0             | 7.4     | 610           | 7.5     | 610           | 7.65    | 610           | 7.9     | 611           | 8.2     | 611           | 8.45    | 611           | 8.9     | 611           | 9.3     | 611           | 9.75    | 611           | 9.9     | 611           | 10.1    | 611           | 10.3    | 611           |
| 7     | 120  | 6.9     | 0             | 7.8     | 916           | 8.6     | 916           | 9.45    | 916           | 10.0    | 917           | 10.6    | 917           | 11.15   | 917           | 11.5    | 917           | 11.8    | 917           | 12.05   | 917           | 12.4    | 917           | 12.7    | 917           | 12.95   | 917           |
| 7     | 125  | 8.85    | 0             | 9.3     | 916           | 9.7     | 916           | 10.05   | 916           | 9.8     | 917           | 9.6     | 917           | 9.4     | 917           | 10.4    | 917           | 11.4    | 917           | 12.35   | 917           | 12.6    | 917           | 12.9    | 917           | 13.15   | 917           |
| 7     | 138  | 8.6     | 0             | 8.7     | 916           | 8.8     | 916           | 8.9     | 916           | 9.0     | 917           | 9.0     | 917           | 9.05    | 917           | 9.7     | 917           | 10.4    | 917           | 11.05   | 917           | 11.6    | 917           | 12.1    | 917           | 12.55   | 917           |
| 7     | 150  | 7.75    | 0             | 7.9     | 916           | 8.1     | 916           | 8.3     | 916           | 8.9     | 917           | 9.5     | 917           | 10.15   | 917           | 10.2    | 917           | 10.3    | 917           | 10.4    | 917           | 10.6    | 917           | 10.8    | 917           | 11      | 917           |
| 8     | 102  | 8.75    | 0             | 8.9     | 300           | 9.1     | 300           | 9.3     | 300           | 10.2    | 300           | 11.0    | 300           | 11.85   | 300           | 11.5    | 300           | 11.1    | 300           | 10.7    | 300           | 11.0    | 300           | 11.4    | 300           | 11.7    | 300           |
| 8     | 114  | 10.1    | 0             | 10.4    | 300           | 10.6    | 300           | 10.9    | 300           | 11.3    | 300           | 11.8    | 300           | 12.2    | 300           | 12.8    | 300           | 13.5    | 300           | 14.1    | 300           | 14.3    | 300           | 14.6    | 300           | 14.8    | 300           |
| 8     | 117  | 10.5    | 0             | 10.7    | 300           | 11.0    | 300           | 11.2    | 300           | 11.5    | 300           | 11.7    | 300           | 11.95   | 300           | 12.5    | 300           | 13.1    | 300           | 13.65   | 300           | 13.9    | 300           | 14.1    | 300           | 14.3    | 300           |
| 8     | 126  | 10.65   | 0             | 10.7    | 300           | 10.8    | 300           | 10.9    | 300           | 10.9    | 300           | 10.8    | 300           | 10.75   | 300           | 11.9    | 300           | 13.0    | 300           | 14.15   | 300           | 14.3    | 300           | 14.5    | 300           | 14.65   | 300           |
| 9     | 112  | 8.75    | 0             | 8.9     | 600           | 9.1     | 600           | 9.3     | 600           | 10.2    | 600           | 11.0    | 600           | 11.85   | 600           | 11.5    | 600           | 11.1    | 600           | 10.7    | 600           | 11.0    | 600           | 11.4    | 600           | 11.7    | 600           |
| 9     | 113  | 10.1    | 0             | 10.4    | 600           | 10.6    | 600           | 10.9    | 600           | 11.3    | 600           | 11.8    | 600           | 12.2    | 600           | 12.8    | 600           | 13.5    | 600           | 14.1    | 600           | 14.3    | 600           | 14.6    | 600           | 14.8    | 600           |
| 9     | 135  | 10.5    | 0             | 10.7    | 600           | 11.0    | 600           | 11.2    | 600           | 11.5    | 600           | 11.7    | 600           | 11.95   | 600           | 12.5    | 600           | 13.1    | 600           | 13.65   | 600           | 13.9    | 600           | 14.1    | 600           | 14.3    | 600           |
| 9     | 154  | 10.65   | 0             | 10.7    | 600           | 10.8    | 600           | 10.9    | 600           | 10.9    | 600           | 10.8    | 600           | 10.75   | 600           | 11.9    | 600           | 13.0    | 600           | 14.15   | 600           | 14.3    | 600           | 14.5    | 600           | 14.65   | 600           |
| 10    | 124  | 8.5     | 0             | 9.0     | 900           | 9.5     | 900           | 10      | 900           | 9.7     | 900           | 9.5     | 900           | 9.21    | 900           | 10.5    | 900           | 11.8    | 900           | 13.1    | 900           | 13.3    | 900           | 13.4    | 900           | 13.55   | 900           |
| 10    | 133  | 7.15    | 0             | 7.4     | 900           | 7.6     | 900           | 7.8     | 900           | 8.3     | 900           | 8.8     | 900           | 9.25    | 900           | 9.6     | 900           | 9.9     | 900           | 10.2    | 900           | 10.5    | 900           | 10.8    | 900           | 11.15   | 900           |
| 10    | 158  | 8.55    | 0             | 8.8     | 900           | 9.0     | 900           | 9.25    | 900           | 9.3     | 900           | 9.3     | 900           | 9.35    | 900           | 10.1    | 900           | 10.9    | 900           | 11.6    | 900           | 11.9    | 900           | 12.2    | 900           | 12.5    | 900           |
| 10    | 160  | 7.65    | 0             | 7.8     | 900           | 7.9     | 900           | 8       | 900           | 8.2     | 900           | 8.4     | 900           | 8.55    | 900           | 8.9     | 900           | 9.3     | 900           | 9.7     | 900           | 10.2    | 900           | 10.7    | 900           | 11.2    | 900           |

**TABLE A-4 BODY WEIGHT ADJUSTED DOSES (ug/kg-day)**

(Dose for Day/BW for Day)

| Group | Pig # | Day 0  | Day 1  | Day 2  | Day 3  | Day 4  | Day 5  | Day 6  | Day 7 | Day 8 | Day 9 | Day 10 | Day 11 | Avg Dose | Avg Dose per Group |
|-------|-------|--------|--------|--------|--------|--------|--------|--------|-------|-------|-------|--------|--------|----------|--------------------|
| 1     | 108   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00  | 0.00  | 0.00  | 0.00   | 0.00   | 0.00     | 0                  |
| 1     | 145   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00  | 0.00  | 0.00  | 0.00   | 0.00   | 0.00     |                    |
| 1     | 157   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00   | 0.00  | 0.00  | 0.00  | 0.00   | 0.00   | 0.00     |                    |
| 2     | 122   | 35.79  | 34.62  | 33.52  | 32.20  | 30.98  | 29.85  | 31.86  | 34.16 | 36.81 | 31.25 | 27.15  | 24.00  | 31.85    | 33.63              |
| 2     | 123   | 39.56  | 37.89  | 36.36  | 34.75  | 33.27  | 31.91  | 30.87  | 29.90 | 28.99 | 28.13 | 27.31  | 26.55  | 32.13    |                    |
| 2     | 147   | 37.74  | 37.27  | 36.81  | 35.86  | 34.95  | 34.09  | 34.03  | 33.96 | 33.90 | 32.26 | 30.77  | 29.41  | 34.25    |                    |
| 2     | 156   | 39.91  | 38.79  | 37.74  | 37.19  | 36.66  | 36.14  | 36.29  | 36.44 | 36.59 | 34.82 | 33.21  | 31.75  | 36.29    |                    |
| 3     | 101   | 71.15  | 65.81  | 61.22  | 58.92  | 56.78  | 54.79  | 53.81  | 52.86 | 51.95 | 49.38 | 47.06  | 44.94  | 55.72    | 59.05              |
| 3     | 115   | 75.16  | 71.29  | 67.80  | 65.45  | 63.27  | 61.22  | 60.81  | 60.40 | 60.00 | 56.69 | 53.73  | 51.06  | 62.24    |                    |
| 3     | 119   | 64.75  | 62.94  | 61.22  | 59.80  | 58.44  | 57.14  | 56.78  | 56.43 | 56.07 | 52.40 | 49.18  | 46.33  | 56.79    |                    |
| 3     | 151   | 71.57  | 69.23  | 67.04  | 65.81  | 64.63  | 63.49  | 60.50  | 57.78 | 55.30 | 54.63 | 53.97  | 53.33  | 61.44    |                    |
| 4     | 121   | 149.17 | 130.12 | 115.38 | 112.97 | 110.66 | 108.43 | 100.37 | 93.43 | 87.38 | 85.85 | 84.38  | 82.95  | 105.09   | 93.66              |
| 4     | 136   | 94.90  | 93.59  | 92.31  | 88.67  | 85.31  | 82.19  | 80.60  | 79.06 | 77.59 | 75.00 | 72.58  | 70.31  | 82.68    |                    |
| 4     | 140   | 119.21 | 110.43 | 102.86 | 100.37 | 98.00  | 95.74  | 95.74  | 95.74 | 95.74 | 90.00 | 84.91  | 80.36  | 97.43    |                    |
| 4     | 148   | 109.76 | 104.05 | 98.90  | 95.58  | 92.47  | 89.55  | 86.68  | 83.98 | 81.45 | 79.18 | 77.03  | 75.00  | 89.47    |                    |
| 5     | 104   | 34.88  | 34.10  | 33.36  | 32.05  | 30.81  | 29.66  | 29.66  | 29.66 | 29.66 | 27.99 | 26.49  | 25.15  | 30.29    | 29.92              |
| 5     | 106   | 35.22  | 34.95  | 34.68  | 33.39  | 32.16  | 31.02  | 29.24  | 27.65 | 26.23 | 25.68 | 25.15  | 24.64  | 30.00    |                    |
| 5     | 128   | 32.53  | 30.78  | 29.21  | 28.42  | 27.65  | 26.92  | 27.78  | 28.69 | 29.66 | 27.53 | 25.68  | 24.06  | 28.24    |                    |
| 5     | 155   | 39.30  | 37.30  | 35.49  | 34.27  | 33.09  | 31.99  | 29.91  | 28.07 | 26.45 | 26.23 | 26.00  | 25.78  | 31.16    |                    |
| 6     | 103   | 80.50  | 78.09  | 75.83  | 72.46  | 69.31  | 66.42  | 63.22  | 60.30 | 57.65 | 56.06 | 54.56  | 53.14  | 65.63    | 60.18              |
| 6     | 110   | 66.96  | 62.72  | 58.98  | 57.02  | 55.14  | 53.37  | 51.14  | 49.08 | 47.19 | 46.00 | 44.88  | 43.81  | 53.02    |                    |
| 6     | 116   | 62.40  | 61.25  | 60.14  | 57.20  | 54.48  | 52.01  | 49.41  | 47.07 | 44.93 | 44.17 | 43.44  | 42.73  | 51.60    |                    |
| 6     | 142   | 82.31  | 81.03  | 79.80  | 77.19  | 74.67  | 72.32  | 68.79  | 65.59 | 62.68 | 61.52 | 60.40  | 59.33  | 70.47    |                    |
| 7     | 120   | 118.15 | 106.47 | 96.89  | 91.51  | 86.61  | 82.21  | 80.06  | 78.01 | 76.07 | 74.22 | 72.46  | 70.78  | 86.12    | 90.04              |
| 7     | 125   | 98.99  | 94.89  | 91.11  | 93.22  | 95.32  | 97.51  | 88.28  | 80.64 | 74.22 | 72.65 | 71.15  | 69.71  | 85.64    |                    |
| 7     | 138   | 105.25 | 104.05 | 102.88 | 102.42 | 101.85 | 101.29 | 94.34  | 88.28 | 82.95 | 79.36 | 76.07  | 73.04  | 92.65    |                    |
| 7     | 150   | 115.42 | 112.81 | 110.32 | 102.80 | 96.15  | 90.31  | 89.57  | 88.85 | 88.14 | 86.47 | 84.87  | 83.33  | 95.75    |                    |
| 8     | 102   | 33.58  | 32.91  | 32.26  | 29.56  | 27.27  | 25.32  | 26.16  | 27.07 | 28.04 | 27.19 | 26.39  | 25.64  | 28.45    | 25.44              |
| 8     | 114   | 28.94  | 28.21  | 27.52  | 26.47  | 25.50  | 24.59  | 23.38  | 22.28 | 21.28 | 20.93 | 20.59  | 20.27  | 24.16    |                    |
| 8     | 117   | 27.95  | 27.36  | 26.79  | 26.20  | 25.64  | 25.10  | 23.97  | 22.93 | 21.98 | 21.63 | 21.30  | 20.98  | 24.32    |                    |
| 8     | 126   | 27.95  | 27.73  | 27.52  | 27.65  | 27.78  | 27.91  | 25.25  | 23.05 | 21.20 | 20.95 | 20.71  | 20.48  | 24.85    |                    |
| 9     | 112   | 67.16  | 65.81  | 64.52  | 59.11  | 54.55  | 50.63  | 52.33  | 54.14 | 56.07 | 54.38 | 52.79  | 51.28  | 56.90    | 50.89              |
| 9     | 113   | 57.88  | 56.43  | 55.05  | 52.94  | 50.99  | 49.18  | 46.75  | 44.55 | 42.55 | 41.86 | 41.19  | 40.54  | 48.33    |                    |
| 9     | 135   | 55.90  | 54.71  | 53.57  | 52.40  | 51.28  | 50.21  | 47.94  | 45.86 | 43.96 | 43.27 | 42.60  | 41.96  | 48.64    |                    |
| 9     | 154   | 55.90  | 55.47  | 55.05  | 55.30  | 55.56  | 55.81  | 50.49  | 46.09 | 42.40 | 41.91 | 41.43  | 40.96  | 49.70    |                    |
| 10    | 124   | 100.00 | 94.74  | 90.00  | 92.43  | 95.00  | 97.72  | 85.66  | 76.25 | 68.70 | 67.92 | 67.16  | 66.42  | 83.50    | 92.88              |
| 10    | 133   | 122.17 | 118.68 | 115.38 | 108.65 | 102.66 | 97.30  | 94.08  | 91.06 | 88.24 | 85.58 | 83.08  | 80.72  | 98.97    |                    |
| 10    | 158   | 102.47 | 99.82  | 97.30  | 96.95  | 96.60  | 96.26  | 89.11  | 82.95 | 77.59 | 75.63 | 73.77  | 72.00  | 88.37    |                    |
| 10    | 160   | 115.88 | 114.16 | 112.50 | 109.98 | 107.57 | 105.26 | 100.75 | 96.60 | 92.78 | 88.24 | 84.11  | 80.36  | 100.68   |                    |

**TABLE A-5 URINE VOLUMES - 48 HOUR COLLECTIONS**

Units of Volume: mL

| Group | Pig # | Day              |                  |                   |
|-------|-------|------------------|------------------|-------------------|
|       |       | 6-7<br>6/26-6/28 | 8-9<br>6/28-6/30 | 10-11<br>6/30-7/2 |
| 1     | 108   | 9580             | 4800             | 13600             |
|       | 145   | 4640             | 5000             | 5800              |
|       | 157   | 10100            | 7500             | 6400              |
| 2     | 122   | 6980             | 7000             | 6200              |
|       | 123   | 24780            | 28600            | 28200             |
|       | 147   | 4880             | 9000             | 5800              |
|       | 156   | 5120             | 4800             | 4500              |
| 3     | 101   | 14360            | 3470             | 59500             |
|       | 115   | 35820            | 35600            | 35200             |
|       | 119   | 3000             | 3800             | 2780              |
|       | 151   | 3620             | 4500             | 6500              |
| 4     | 121   | 8420             | 1400             | 16000             |
|       | 136   | 2364             | 14800            | 14200             |
|       | 140   | 9840             | 11800            | 14840             |
|       | 148   | 12360            | 14900            | 10200             |
| 5     | 104   | 3700             | 6400             | 2700              |
|       | 106   | 3720             | 6200             | 2400              |
|       | 128   | 7820             | 10600            | 5000              |
|       | 155   | 3920             | 5600             | 3200              |
| 6     | 103   | 9740             | 9700             | 4200              |
|       | 110   | 8100             | 8600             | 3200              |
|       | 116   | 14000            | 15600            | 6500              |
|       | 142   | 14000            | 7800             | 3100              |
| 7     | 120   | 5720             | 9800             | 3500              |
|       | 125   | 15590            | 21200            | 14600             |
|       | 138   | 6960             | 3100             | 1200              |
|       | 150   | 10200            | 8400             | 6000              |
| 8     | 102   | 4640             | 6200             | 5200              |
|       | 114   | 8520             | 11000            | 9200              |
|       | 117   | 10560            | 11000            | 17800             |
|       | 126   | 20640            | 29600            | 42860             |
| 9     | 112   | 8700             | 11400            | 13000             |
|       | 113   | 11040            | 1600             | 16600             |
|       | 135   | 6940             | 9400             | 14000             |
|       | 154   | 2300             | 7800             | 6800              |
| 10    | 124   | 3800             | 5400             | 6400              |
|       | 133   | 7200             | 7200             | 8600              |
|       | 158   | 1940             | 4800             | 4300              |
|       | 160   | 7240             | 7800             | 7600              |

= Volumes are for a 24-hour collection on Day 10.

**TABLE A-6 FECAL WEIGHTS - 48 HOUR COLLECTIONS**

Units of Weight:      grams

| Group | Pig ID | Day              |                  |                   |
|-------|--------|------------------|------------------|-------------------|
|       |        | 6-7<br>6/26-6/28 | 8-9<br>6/28-6/30 | 10-11<br>6/30-7/2 |
| 1     | 108    | 284.8            | 290.9            | 363.7             |
|       | 145    | 265.6            | 240.2            | 305.8             |
|       | 157    | 225.9            | 395.9            | 4568.9            |
| 2     | 122    | 301.1            | 340.8            | 296.2             |
|       | 123    | 264.9            | 243              | 311.3             |
|       | 147    | 228.6            | 373.5            | 171.4             |
|       | 156    | missing          | 86.2             | 259.8             |
| 3     | 101    | 372.8            | 303.2            | 376.2             |
|       | 115    | 321.9            | 305.8            | 305.9             |
|       | 119    | 300.3            | 569.7            | 360.6             |
|       | 151    | 289              | 465              | 380.7             |
| 4     | 121    | 381.4            | 392.8            | 347               |
|       | 136    | 522.5            | 342              | 365.7             |
|       | 140    | 31.9             | 414              | 212.4             |
|       | 148    | 227.4            | 316.7            | 331               |
| 5     | 104    | 317.7            | 411.1            | 366.4             |
|       | 106    | 236.3            | 558.3            | 213.2             |
|       | 128    | 359.7            | 354.4            | 218.1             |
|       | 155    | 436.3            | 356              | 145.1             |
| 6     | 103    | 211.3            | 412.4            | 291.8             |
|       | 110    | 265.1            | 526              | 218.9             |
|       | 116    | 361.8            | 652              | 356.7             |
|       | 142    | 276.1            | 401.1            | 243.5             |
| 7     | 120    | 296.8            | 432              | 204.1             |
|       | 125    | 373.4            | 428.1            | 264.3             |
|       | 138    | 640.3            | 381.7            | 268.2             |
|       | 150    | 333.9            | 405.8            | 253.5             |
| 8     | 102    | 337.9            | 438.5            | 240.9             |
|       | 114    | 361.8            | 403.2            | 244.2             |
|       | 117    | 387.1            | 376.5            | 198               |
|       | 126    | 543.2            | 489.3            | 363.2             |
| 9     | 112    | 582.8            | 541.5            | 275               |
|       | 113    | 602.9            | 413.3            | 384.5             |
|       | 135    | 443.8            | 364              | 165.1             |
|       | 154    | 498.1            | 467.3            | 372.6             |
| 10    | 124    | 490.2            | 341.2            | 368.9             |
|       | 133    | 483.9            | 506.6            | 190.7             |
|       | 158    | 534              | 265.1            | 369.5             |
|       | 160    | 192.6            | 327              | 268.3             |

= Weights are for a 24-hour collection on Day 10.

**TABLE A-7 URINE ANALYTICAL RESULTS**

| ID | pig number | group | dosage | day | date collected | sample number       | tag number2 | MgNO3 Q | MgNO3 ng/mL |
|----|------------|-------|--------|-----|----------------|---------------------|-------------|---------|-------------|
| 1  | 108        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-108-(6/7)-U     | BA-01-00205 |         | 2           |
| 2  | 145        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-145-(6/7)-U     | BA-01-00209 |         | 2           |
| 3  | 157        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-157-(6/7)-U     | BA-01-00232 |         | 2           |
| 4  | 122        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-122-(6/7)-U     | BA-01-00238 |         | 82          |
| 5  | 123        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-123-(6/7)-U     | BA-01-00187 |         | 20          |
| 6  | 147        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-147-(6/7)-U     | BA-01-00192 |         | 68          |
| 7  | 156        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-156-(6/7)-U     | BA-01-00222 |         | 73          |
| 8  | 101        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-101-(6/7)-U     | BA-01-00190 |         | 71          |
| 9  | 115        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-115-(6/7)-U     | BA-01-00195 |         | 30          |
| 10 | 119        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-119-(6/7)-U     | BA-01-00210 |         | 370         |
| 11 | 151        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-151-(6/7)-U     | BA-01-00233 |         | 300         |
| 12 | 121        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-121-(6/7)-U     | BA-01-00188 |         | 370         |
| 13 | 136        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-136-(6/7)-U     | BA-01-00189 |         | 71          |
| 14 | 140        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-140-(6/7)-U     | BA-01-00213 |         | 150         |
| 15 | 148        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-148-(6/7)-U     | BA-01-00204 |         | 180         |
| 16 | 104        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-104-(6/7)-U     | BA-01-00208 |         | 23          |
| 17 | 106        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-106-(6/7)-U     | BA-01-00198 |         | 19          |
| 18 | 128        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-128-(6/7)-U     | BA-01-00186 |         | 13          |
| 19 | 155        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-155-(6/7)-U     | BA-01-00203 |         | 23          |
| 20 | 103        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-103-(6/7)-U     | BA-01-00217 |         | 16          |
| 21 | 110        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-110-(6/7)-U     | BA-01-00184 |         | 22          |
| 22 | 116        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-116-(6/7)-U     | BA-01-00215 |         | 15          |
| 23 | 142        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-142-(6/7)-U     | BA-01-00219 |         | 12          |
| 24 | 120        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-120-(6/7)-U     | BA-01-00197 |         | 41          |
| 25 | 125        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-125-(6/7)-U     | BA-01-00220 |         | 24          |
| 26 | 138        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-138-(6/7)-U     | BA-01-00236 |         | 35          |
| 27 | 150        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-150-(6/7)-U     | BA-01-00216 |         | 25          |
| 28 | 102        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-102-(6/7)-U     | BA-01-00201 |         | 19          |
| 29 | 114        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-114-(6/7)-U     | BA-01-00234 |         | 15          |
| 30 | 117        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-117-(6/7)-U     | BA-01-00225 |         | 12          |
| 31 | 126        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-126-(6/7)-U     | BA-01-00224 |         | 5.4         |
| 32 | 112        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-112-(6/7)-U     | BA-01-00200 |         | 35          |
| 33 | 113        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-113-(6/7)-U     | BA-01-00191 |         | 19          |
| 34 | 135        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-135-(6/7)-U     | BA-01-00196 |         | 18          |
| 35 | 154        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-154-(6/7)-U     | BA-01-00230 |         | 67          |
| 36 | 124        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-124-(6/7)-U     | BA-01-00207 |         | 96          |
| 37 | 133        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-133-(6/7)-U     | BA-01-00235 |         | 60          |
| 38 | 158        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-158-(6/7)-U     | BA-01-00199 |         | 140         |
| 39 | 160        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-160-(6/7)-U     | BA-01-00218 |         | 69          |
| 40 | 2108       | 1     | 0      | 6/7 | 26-Jun-01      | BA1-2108-(6/7)-U    | BA-01-00211 |         | 2           |
| 41 | 2125       | 7     | 75     | 6/7 | 26-Jun-01      | BA1-2125-(6/7)-U    | BA-01-00231 |         | 23          |
| 42 | 2115       | 3     | 50     | 6/7 | 26-Jun-01      | BA1-2115-(6/7)-U    | BA-01-00185 |         | 29          |
| 43 | AsCtrl     |       |        | 6/7 | 26-Jun-01      | BA1-AsCtrl-(6/7)-U  | BA-01-00194 |         | 3           |
| 44 | AsIA500    |       |        | 6/7 | 26-Jun-01      | BA1-AsIA500-(6/7)-U | BA-01-00229 |         | 520         |
| 45 | AsIB500    |       |        | 6/7 | 26-Jun-01      | BA1-AsIB500-(6/7)-U | BA-01-00214 |         | 500         |
| 46 | AsOA500    |       |        | 6/7 | 26-Jun-01      | BA1-AsOA500-(6/7)-U | BA-01-00226 |         | 510         |
| 47 | AsOB500    |       |        | 6/7 | 26-Jun-01      | BA1-AsOB500-(6/7)-U | BA-01-00227 |         | 510         |
| 48 | AsIA100    |       |        | 6/7 | 26-Jun-01      | BA1-AsIA100-(6/7)-U | BA-01-00206 |         | 100         |
| 49 | AsIB100    |       |        | 6/7 | 26-Jun-01      | BA1-AsIB100-(6/7)-U | BA-01-00221 |         | 100         |
| 50 | AsOA100    |       |        | 6/7 | 26-Jun-01      | BA1-AsOA100-(6/7)-U | BA-01-00237 |         | 100         |
| 51 | AsOB100    |       |        | 6/7 | 26-Jun-01      | BA1-AsOB100-(6/7)-U | BA-01-00228 |         | 100         |
| 52 | AsIA25     |       |        | 6/7 | 26-Jun-01      | BA1-AsIA25-(6/7)-U  | BA-01-00212 |         | 28          |
| 53 | AsIB25     |       |        | 6/7 | 26-Jun-01      | BA1-AsIB25-(6/7)-U  | BA-01-00223 |         | 27          |

**TABLE A-7 URINE ANALYTICAL RESULTS**

| ID  | pig number | group | dosage | day | date collected | sample number       | tag number2 | MgNO3 Q | MgNO3 ng/mL |
|-----|------------|-------|--------|-----|----------------|---------------------|-------------|---------|-------------|
| 54  | AsOA25     |       |        | 6/7 | 26-Jun-01      | BA1-AsOA25-(6/7)-U  | BA-01-00202 |         | 28          |
| 55  | AsOB25     |       |        | 6/7 | 26-Jun-01      | BA1-AsOB25-(6/7)-U  | BA-01-00193 |         | 28          |
| 56  | 108        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-108-(8/9)-U     | BA-01-00288 |         | 2           |
| 57  | 145        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-145-(8/9)-U     | BA-01-00281 |         | 3           |
| 58  | 157        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-157-(8/9)-U     | BA-01-00265 |         | 2           |
| 59  | 122        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-122-(8/9)-U     | BA-01-00243 |         | 82          |
| 60  | 123        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-123-(8/9)-U     | BA-01-00273 |         | 18          |
| 61  | 147        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-147-(8/9)-U     | BA-01-00291 |         | 51          |
| 62  | 156        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-156-(8/9)-U     | BA-01-00269 |         | 80          |
| 63  | 101        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-101-(8/9)-U     | BA-01-00245 |         | 31          |
| 64  | 115        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-115-(8/9)-U     | BA-01-00257 |         | 37          |
| 65  | 119        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-119-(8/9)-U     | BA-01-00282 |         | 270         |
| 66  | 151        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-151-(8/9)-U     | BA-01-00289 |         | 230         |
| 67  | 121        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-121-(8/9)-U     | BA-01-00292 |         | 130         |
| 68  | 136        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-136-(8/9)-U     | BA-01-00261 |         | 110         |
| 69  | 140        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-140-(8/9)-U     | BA-01-00253 |         | 140         |
| 70  | 148        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-148-(8/9)-U     | BA-01-00247 |         | 130         |
| 71  | 104        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-104-(8/9)-U     | BA-01-00244 |         | 17          |
| 72  | 106        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-106-(8/9)-U     | BA-01-00251 |         | 16          |
| 73  | 128        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-128-(8/9)-U     | BA-01-00240 |         | 14          |
| 74  | 155        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-155-(8/9)-U     | BA-01-00250 |         | 17          |
| 75  | 103        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-103-(8/9)-U     | BA-01-00252 |         | 19          |
| 76  | 110        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-110-(8/9)-U     | BA-01-00239 |         | 28          |
| 77  | 116        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-116-(8/9)-U     | BA-01-00256 |         | 15          |
| 78  | 142        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-142-(8/9)-U     | BA-01-00290 |         | 27          |
| 79  | 120        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-120-(8/9)-U     | BA-01-00276 |         | missing     |
| 80  | 125        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-125-(8/9)-U     | BA-01-00258 |         | 19          |
| 81  | 138        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-138-(8/9)-U     | BA-01-00268 |         | 84          |
| 82  | 150        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-150-(8/9)-U     | BA-01-00286 |         | 34          |
| 83  | 102        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-102-(8/9)-U     | BA-01-00293 |         | 15          |
| 84  | 114        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-114-(8/9)-U     | BA-01-00241 |         | 15          |
| 85  | 117        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-117-(8/9)-U     | BA-01-00242 |         | 12          |
| 86  | 126        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-126-(8/9)-U     | BA-01-00272 |         | 6.5         |
| 87  | 112        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-112-(8/9)-U     | BA-01-00262 |         | 25          |
| 88  | 113        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-113-(8/9)-U     | BA-01-00267 |         | 18          |
| 89  | 135        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-135-(8/9)-U     | BA-01-00277 |         | 37          |
| 90  | 154        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-154-(8/9)-U     | BA-01-00287 |         | 28          |
| 91  | 124        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-124-(8/9)-U     | BA-01-00271 |         | 80          |
| 92  | 133        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-133-(8/9)-U     | BA-01-00283 |         | 58          |
| 93  | 158        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-158-(8/9)-U     | BA-01-00259 |         | 62          |
| 94  | 160        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-160-(8/9)-U     | BA-01-00270 |         | 57          |
| 95  | 2135       | 9     | 50     | 8/9 | 28-Jun-01      | BA1-2135-(8/9)-U    | BA-01-00249 |         | 35          |
| 96  | 2103       | 6     | 50     | 8/9 | 28-Jun-01      | BA1-2103-(8/9)-U    | BA-01-00248 |         | 17          |
| 97  | 2157       | 1     | 0      | 8/9 | 28-Jun-01      | BA1-2157-(8/9)-U    | BA-01-00274 |         | 2           |
| 98  | AsCtrl     |       |        | 8/9 | 28-Jun-01      | BA1-AsCtrl-(8/9)-U  | BA-01-00280 |         | 3           |
| 99  | AsIA500    |       |        | 8/9 | 28-Jun-01      | BA1-AsIA500-(8/9)-U | BA-01-00284 |         | 560         |
| 100 | AsIB500    |       |        | 8/9 | 28-Jun-01      | BA1-AsIB500-(8/9)-U | BA-01-00254 |         | 460         |
| 101 | AsOA500    |       |        | 8/9 | 28-Jun-01      | BA1-AsOA500-(8/9)-U | BA-01-00278 |         | 510         |
| 102 | AsOB500    |       |        | 8/9 | 28-Jun-01      | BA1-AsOB500-(8/9)-U | BA-01-00279 |         | 480         |
| 103 | AsIA100    |       |        | 8/9 | 28-Jun-01      | BA1-AsIA100-(8/9)-U | BA-01-00266 |         | 100         |
| 104 | AsIB100    |       |        | 8/9 | 28-Jun-01      | BA1-AsIB100-(8/9)-U | BA-01-00285 |         | 110         |
| 105 | AsOA100    |       |        | 8/9 | 28-Jun-01      | BA1-AsOA100-(8/9)-U | BA-01-00255 |         | 98          |
| 106 | AsOB100    |       |        | 8/9 | 28-Jun-01      | BA1-AsOB100-(8/9)-U | BA-01-00260 |         | 99          |

**TABLE A-7 URINE ANALYTICAL RESULTS**

| ID  | pig number | group | dosage | day   | date collected | sample number        | tag number2 | MgNO3 Q | MgNO3 ng/mL |
|-----|------------|-------|--------|-------|----------------|----------------------|-------------|---------|-------------|
| 107 | AsIA25     |       |        | 8/9   | 28-Jun-01      | BA1-AsIA25-(8/9)-U   | BA-01-00264 |         | 28          |
| 108 | AsIB25     |       |        | 8/9   | 28-Jun-01      | BA1-AsIB25-(8/9)-U   | BA-01-00246 |         | 27          |
| 109 | AsOA25     |       |        | 8/9   | 28-Jun-01      | BA1-AsOA25-(8/9)-U   | BA-01-00275 |         | 28          |
| 110 | AsOB25     |       |        | 8/9   | 28-Jun-01      | BA1-AsOB25-(8/9)-U   | BA-01-00263 |         | 26          |
| 111 | 108        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-108-(10/11)-U    | BA-01-00326 |         | 1.8         |
| 112 | 145        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-145-(10/11)-U    | BA-01-00314 |         | 2           |
| 113 | 157        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-157-(10/11)-U    | BA-01-00297 |         | 2           |
| 114 | 122        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-122-(10/11)-U    | BA-01-00300 |         | 89          |
| 115 | 123        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-123-(10/11)-U    | BA-01-00328 |         | 20          |
| 116 | 147        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-147-(10/11)-U    | BA-01-00334 |         | 80          |
| 117 | 156        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-156-(10/11)-U    | BA-01-00309 |         | 87          |
| 118 | 101        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-101-(10/11)-U    | BA-01-00335 |         | 21          |
| 119 | 115        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-115-(10/11)-U    | BA-01-00304 |         | 36          |
| 120 | 119        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-119-(10/11)-U    | BA-01-00332 |         | 400         |
| 121 | 151        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-151-(10/11)-U    | BA-01-00305 |         | 190         |
| 122 | 121        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-121-(10/11)-U    | BA-01-00322 |         | 140         |
| 123 | 136        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-136-(10/11)-U    | BA-01-00321 |         | 120         |
| 124 | 140        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-140-(10/11)-U    | BA-01-00306 |         | 110         |
| 125 | 148        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-148-(10/11)-U    | BA-01-00311 |         | 140         |
| 126 | 104        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-104-(10/11)-U    | BA-01-00316 |         | 19          |
| 127 | 106        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-106-(10/11)-U    | BA-01-00303 |         | 23          |
| 128 | 128        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-128-(10/11)-U    | BA-01-00294 |         | missing     |
| 129 | 155        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-155-(10/11)-U    | BA-01-00313 |         | 20          |
| 130 | 103        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-103-(10/11)-U    | BA-01-00301 |         | 21          |
| 131 | 110        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-110-(10/11)-U    | BA-01-00318 |         | 32          |
| 132 | 116        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-116-(10/11)-U    | BA-01-00329 |         | 17          |
| 133 | 142        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-142-(10/11)-U    | BA-01-00298 |         | 35          |
| 134 | 120        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-120-(10/11)-U    | BA-01-00330 |         | 45          |
| 135 | 125        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-125-(10/11)-U    | BA-01-00308 |         | 15          |
| 136 | 138        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-138-(10/11)-U    | BA-01-00315 |         | 120         |
| 137 | 150        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-150-(10/11)-U    | BA-01-00319 |         | 23          |
| 138 | 102        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-102-(10/11)-U    | BA-01-00320 |         | 16          |
| 139 | 114        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-114-(10/11)-U    | BA-01-00317 |         | 19          |
| 140 | 117        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-117-(10/11)-U    | BA-01-00302 |         | 7.7         |
| 141 | 126        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-126-(10/11)-U    | BA-01-00333 |         | 5.4         |
| 142 | 112        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-112-(10/11)-U    | BA-01-00325 |         | 25          |
| 143 | 113        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-113-(10/11)-U    | BA-01-00323 |         | 19          |
| 144 | 135        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-135-(10/11)-U    | BA-01-00324 |         | 5.9         |
| 145 | 154        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-154-(10/11)-U    | BA-01-00299 |         | 40          |
| 146 | 124        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-124-(10/11)-U    | BA-01-00312 |         | 33          |
| 147 | 133        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-133-(10/11)-U    | BA-01-00296 |         | 49          |
| 148 | 158        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-158-(10/11)-U    | BA-01-00327 |         | 96          |
| 149 | 160        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-160-(10/11)-U    | BA-01-00295 |         | 47          |
| 150 | 2145       | 1     | 0      | 10/11 | 30-Jun-01      | BA1-2145-(10/11)-U   | BA-01-00310 |         | 1           |
| 151 | 2133       | 10    | 75     | 10/11 | 30-Jun-01      | BA1-2133-(10/11)-U   | BA-01-00307 |         | 48          |
| 152 | 2148       | 4     | 75     | 10/11 | 30-Jun-01      | BA1-2148-(10/11)-U   | BA-01-00336 |         | 140         |
| 153 | AsCtrl     |       |        | 10/11 | 30-Jun-01      | BA1-AsCtrl-(10/11)-U | BA-01-00331 |         | 3           |

**TABLE A-8 FECES ANALYTICAL RESULTS**

| ID | pig number | group | dosage | day | date collected | sample number          | tag number  | Q | Conc    |
|----|------------|-------|--------|-----|----------------|------------------------|-------------|---|---------|
| 1  | 108        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-108-(6/7)-P        | BA-01-00346 |   | 30      |
| 2  | 145        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-145-(6/7)-P        | BA-01-00348 |   | 10      |
| 3  | 157        | 1     | 0      | 6/7 | 26-Jun-01      | BA1-157-(6/7)-P        | BA-01-00372 |   | 10      |
| 4  | 122        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-122-(6/7)-P        | BA-01-00362 |   | 73      |
| 5  | 123        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-123-(6/7)-P        | BA-01-00383 |   | 190     |
| 6  | 147        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-147-(6/7)-P        | BA-01-00388 |   | 240     |
| 7  | 156        | 2     | 25     | 6/7 | 26-Jun-01      | BA1-156-(6/7)-P        | BA-01-00341 |   | 540     |
| 8  | 101        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-101-(6/7)-P        | BA-01-00364 |   | 130     |
| 9  | 115        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-115-(6/7)-P        | BA-01-00338 |   | 460     |
| 10 | 119        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-119-(6/7)-P        | BA-01-00350 |   | 2880    |
| 11 | 151        | 3     | 50     | 6/7 | 26-Jun-01      | BA1-151-(6/7)-P        | BA-01-00355 |   | 210     |
| 12 | 121        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-121-(6/7)-P        | BA-01-00387 |   | 310     |
| 13 | 136        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-136-(6/7)-P        | BA-01-00389 |   | 510     |
| 14 | 140        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-140-(6/7)-P        | BA-01-00384 |   | 1400    |
| 15 | 148        | 4     | 75     | 6/7 | 26-Jun-01      | BA1-148-(6/7)-P        | BA-01-00385 |   | 750     |
| 16 | 104        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-104-(6/7)-P        | BA-01-00353 |   | 1400    |
| 17 | 106        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-106-(6/7)-P        | BA-01-00386 |   | 900     |
| 18 | 128        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-128-(6/7)-P        | BA-01-00381 |   | 1000    |
| 19 | 155        | 5     | 25     | 6/7 | 26-Jun-01      | BA1-155-(6/7)-P        | BA-01-00375 |   | 1100    |
| 20 | 103        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-103-(6/7)-P        | BA-01-00347 |   | 2600    |
| 21 | 110        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-110-(6/7)-P        | BA-01-00390 |   | 1500    |
| 22 | 116        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-116-(6/7)-P        | BA-01-00366 |   | 3400    |
| 23 | 142        | 6     | 50     | 6/7 | 26-Jun-01      | BA1-142-(6/7)-P        | BA-01-00342 |   | 3800    |
| 24 | 120        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-120-(6/7)-P        | BA-01-00376 |   | 2000    |
| 25 | 125        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-125-(6/7)-P        | BA-01-00343 |   | 3400    |
| 26 | 138        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-138-(6/7)-P        | BA-01-00365 |   | 5600    |
| 27 | 150        | 7     | 75     | 6/7 | 26-Jun-01      | BA1-150-(6/7)-P        | BA-01-00351 |   | 2700    |
| 28 | 102        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-102-(6/7)-P        | BA-01-00358 |   | 630     |
| 29 | 114        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-114-(6/7)-P        | BA-01-00367 |   | 640     |
| 30 | 117        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-117-(6/7)-P        | BA-01-00356 |   | 1200    |
| 31 | 126        | 8     | 25     | 6/7 | 26-Jun-01      | BA1-126-(6/7)-P        | BA-01-00373 |   | 1200    |
| 32 | 112        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-112-(6/7)-P        | BA-01-00357 |   | 2400    |
| 33 | 113        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-113-(6/7)-P        | BA-01-00349 |   | 2700    |
| 34 | 135        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-135-(6/7)-P        | BA-01-00363 |   | 2200    |
| 35 | 154        | 9     | 50     | 6/7 | 26-Jun-01      | BA1-154-(6/7)-P        | BA-01-00371 |   | 3100    |
| 36 | 124        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-124-(6/7)-P        | BA-01-00352 |   | 3700    |
| 37 | 133        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-133-(6/7)-P        | BA-01-00374 |   | 4800    |
| 38 | 158        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-158-(6/7)-P        | BA-01-00344 |   | 4000    |
| 39 | 160        | 10    | 75     | 6/7 | 26-Jun-01      | BA1-160-(6/7)-P        | BA-01-00339 |   | 1100    |
| 40 | 2142       | 6     | 50     | 6/7 | 26-Jun-01      | BA1-2142-(6/7)-P       | BA-01-00354 |   | 3700    |
| 41 | 2148       | 4     | 75     | 6/7 | 26-Jun-01      | BA1-2148-(6/7)-P       | BA-01-00340 |   | 700     |
| 42 | 2110       | 6     | 50     | 6/7 | 26-Jun-01      | BA1-2110-(6/7)-P       | BA-01-00361 |   | 1900    |
| 43 | FAsCtrl    |       |        | 6/7 | 26-Jun-01      | BA1-FAsCtrl-(6/7)-P    | BA-01-00370 |   | 48      |
| 44 | FAsIAHigh  |       |        | 6/7 | 26-Jun-01      | BA1-FAsIAHigh-(6/7)-P  | BA-01-00379 |   | 3500    |
| 45 | FAsIBHigh  |       |        | 6/7 | 26-Jun-01      | BA1-FAsIBHigh-(6/7)-P  | BA-01-00359 |   | 3200    |
| 46 | FAsOAHHigh |       |        | 6/7 | 26-Jun-01      | BA1-FAsOAHHigh-(6/7)-P | BA-01-00378 |   | 3500    |
| 47 | FAsOBHigh  |       |        | 6/7 | 26-Jun-01      | BA1-FAsOBHigh-(6/7)-P  | BA-01-00337 |   | 2300    |
| 48 | FAsIAMed   |       |        | 6/7 | 26-Jun-01      | BA1-FAsIAMed-(6/7)-P   | BA-01-00368 |   | 1800    |
| 49 | FAsIBMed   |       |        | 6/7 | 26-Jun-01      | BA1-FAsIBMed-(6/7)-P   | BA-01-00345 |   | 1600    |
| 50 | FAsOAMed   |       |        | 6/7 | 26-Jun-01      | BA1-FAsOAMed-(6/7)-P   | BA-01-00380 |   | 1800    |
| 51 | FAsOBMed   |       |        | 6/7 | 26-Jun-01      | BA1-FAsOBMed-(6/7)-P   | BA-01-00369 |   | 1800    |
| 52 | FAsIALow   |       |        | 6/7 | 26-Jun-01      | BA1-FAsIALow-(6/7)-P   | BA-01-00391 |   | 470     |
| 53 | FAsIBLow   |       |        | 6/7 | 26-Jun-01      | BA1-FAsIBLow-(6/7)-P   | BA-01-00360 |   | 430     |
| 54 | FAsOALow   |       |        | 6/7 | 26-Jun-01      | BA1-FAsOALow-(6/7)-P   | BA-01-00382 |   | missing |

**TABLE A-8 FECES ANALYTICAL RESULTS**

| ID  | pig number | group | dosage | day | date collected | sample number          | tag number  | Q | Conc |
|-----|------------|-------|--------|-----|----------------|------------------------|-------------|---|------|
| 55  | FAsOBLow   |       |        | 6/7 | 26-Jun-01      | BA1-FAsOBLow-(6/7)-P   | BA-01-00377 |   | 330  |
| 56  | 108        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-108-(8/9)-P        | BA-01-00446 |   | 81   |
| 57  | 145        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-145-(8/9)-P        | BA-01-00395 |   | 10   |
| 58  | 157        | 1     | 0      | 8/9 | 28-Jun-01      | BA1-157-(8/9)-P        | BA-01-00431 |   | 20   |
| 59  | 122        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-122-(8/9)-P        | BA-01-00422 |   | 59   |
| 60  | 123        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-123-(8/9)-P        | BA-01-00425 |   | 160  |
| 61  | 147        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-147-(8/9)-P        | BA-01-00404 |   | 300  |
| 62  | 156        | 2     | 25     | 8/9 | 28-Jun-01      | BA1-156-(8/9)-P        | BA-01-00415 |   | 75   |
| 63  | 101        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-101-(8/9)-P        | BA-01-00413 |   | 76   |
| 64  | 115        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-115-(8/9)-P        | BA-01-00407 |   | 170  |
| 65  | 119        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-119-(8/9)-P        | BA-01-00445 |   | 150  |
| 66  | 151        | 3     | 50     | 8/9 | 28-Jun-01      | BA1-151-(8/9)-P        | BA-01-00440 |   | 250  |
| 67  | 121        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-121-(8/9)-P        | BA-01-00434 |   | 250  |
| 68  | 136        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-136-(8/9)-P        | BA-01-00409 |   | 340  |
| 69  | 140        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-140-(8/9)-P        | BA-01-00442 |   | 1200 |
| 70  | 148        | 4     | 75     | 8/9 | 28-Jun-01      | BA1-148-(8/9)-P        | BA-01-00443 |   | 320  |
| 71  | 104        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-104-(8/9)-P        | BA-01-00399 |   | 1500 |
| 72  | 106        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-106-(8/9)-P        | BA-01-00444 |   | 800  |
| 73  | 128        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-128-(8/9)-P        | BA-01-00414 |   | 240  |
| 74  | 155        | 5     | 25     | 8/9 | 28-Jun-01      | BA1-155-(8/9)-P        | BA-01-00427 |   | 490  |
| 75  | 103        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-103-(8/9)-P        | BA-01-00428 |   | 2600 |
| 76  | 110        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-110-(8/9)-P        | BA-01-00432 |   | 2500 |
| 77  | 116        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-116-(8/9)-P        | BA-01-00403 |   | 2300 |
| 78  | 142        | 6     | 50     | 8/9 | 28-Jun-01      | BA1-142-(8/9)-P        | BA-01-00437 |   | 3000 |
| 79  | 120        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-120-(8/9)-P        | BA-01-00420 |   | 3300 |
| 80  | 125        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-125-(8/9)-P        | BA-01-00405 |   | 2700 |
| 81  | 138        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-138-(8/9)-P        | BA-01-00417 |   | 3000 |
| 82  | 150        | 7     | 75     | 8/9 | 28-Jun-01      | BA1-150-(8/9)-P        | BA-01-00394 |   | 1900 |
| 83  | 102        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-102-(8/9)-P        | BA-01-00397 |   | 980  |
| 84  | 114        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-114-(8/9)-P        | BA-01-00396 |   | 900  |
| 85  | 117        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-117-(8/9)-P        | BA-01-00392 |   | 1200 |
| 86  | 126        | 8     | 25     | 8/9 | 28-Jun-01      | BA1-126-(8/9)-P        | BA-01-00419 |   | 1200 |
| 87  | 112        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-112-(8/9)-P        | BA-01-00421 |   | 2800 |
| 88  | 113        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-113-(8/9)-P        | BA-01-00402 |   | 770  |
| 89  | 135        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-135-(8/9)-P        | BA-01-00438 |   | 2100 |
| 90  | 154        | 9     | 50     | 8/9 | 28-Jun-01      | BA1-154-(8/9)-P        | BA-01-00430 |   | 1700 |
| 91  | 124        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-124-(8/9)-P        | BA-01-00406 |   | 2800 |
| 92  | 133        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-133-(8/9)-P        | BA-01-00401 |   | 2900 |
| 93  | 158        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-158-(8/9)-P        | BA-01-00439 |   | 3500 |
| 94  | 160        | 10    | 75     | 8/9 | 28-Jun-01      | BA1-160-(8/9)-P        | BA-01-00441 |   | 3600 |
| 95  | 2104       | 5     | 25     | 8/9 | 28-Jun-01      | BA1-2104-(8/9)-P       | BA-01-00429 |   | 1600 |
| 96  | 2136       | 4     | 75     | 8/9 | 28-Jun-01      | BA1-2136-(8/9)-P       | BA-01-00424 |   | 300  |
| 97  | 2148       | 4     | 75     | 8/9 | 28-Jun-01      | BA1-2148-(8/9)-P       | BA-01-00408 |   | 340  |
| 98  | FAsCtrl    |       |        | 8/9 | 28-Jun-01      | BA1-FAsCtrl-(8/9)-P    | BA-01-00412 |   | 29   |
| 99  | FAsIAHigh  |       |        | 8/9 | 28-Jun-01      | BA1-FAsIAHigh-(8/9)-P  | BA-01-00435 |   | 3600 |
| 100 | FAsIBHigh  |       |        | 8/9 | 28-Jun-01      | BA1-FAsIBHigh-(8/9)-P  | BA-01-00398 |   | 3300 |
| 101 | FAsOAHHigh |       |        | 8/9 | 28-Jun-01      | BA1-FAsOAHHigh-(8/9)-P | BA-01-00416 |   | 3400 |
| 102 | FAsOBHigh  |       |        | 8/9 | 28-Jun-01      | BA1-FAsOBHigh-(8/9)-P  | BA-01-00410 |   | 2400 |
| 103 | FAsIAMed   |       |        | 8/9 | 28-Jun-01      | BA1-FAsIAMed-(8/9)-P   | BA-01-00418 |   | 1800 |
| 104 | FAsIBMed   |       |        | 8/9 | 28-Jun-01      | BA1-FAsIBMed-(8/9)-P   | BA-01-00436 |   | 1700 |
| 105 | FAsOAMed   |       |        | 8/9 | 28-Jun-01      | BA1-FAsOAMed-(8/9)-P   | BA-01-00393 |   | 1800 |
| 106 | FAsOBMed   |       |        | 8/9 | 28-Jun-01      | BA1-FAsOBMed-(8/9)-P   | BA-01-00400 |   | 1700 |
| 107 | FAsIALow   |       |        | 8/9 | 28-Jun-01      | BA1-FAsIALow-(8/9)-P   | BA-01-00426 |   | 450  |
| 108 | FAsIBLow   |       |        | 8/9 | 28-Jun-01      | BA1-FAsIBLow-(8/9)-P   | BA-01-00411 |   | 450  |

**TABLE A-8 FECES ANALYTICAL RESULTS**

| ID  | pig number | group | dosage | day   | date collected | sample number         | tag number  | Q | Conc    |
|-----|------------|-------|--------|-------|----------------|-----------------------|-------------|---|---------|
| 109 | FAsOALow   |       |        | 8/9   | 28-Jun-01      | BA1-FAsOALow-(8/9)-P  | BA-01-00423 |   | missing |
| 110 | FAsOBLow   |       |        | 8/9   | 28-Jun-01      | BA1-FAsOBLow-(8/9)-P  | BA-01-00433 |   | 380     |
| 111 | 108        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-108-(10/11)-P     | BA-01-00479 |   | 99      |
| 112 | 145        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-145-(10/11)-P     | BA-01-00456 | < | 10      |
| 113 | 157        | 1     | 0      | 10/11 | 30-Jun-01      | BA1-157-(10/11)-P     | BA-01-00461 |   | 10      |
| 114 | 122        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-122-(10/11)-P     | BA-01-00463 |   | 110     |
| 115 | 123        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-123-(10/11)-P     | BA-01-00452 |   | 100     |
| 116 | 147        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-147-(10/11)-P     | BA-01-00448 |   | 340     |
| 117 | 156        | 2     | 25     | 10/11 | 30-Jun-01      | BA1-156-(10/11)-P     | BA-01-00451 |   | 280     |
| 118 | 101        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-101-(10/11)-P     | BA-01-00454 |   | 85      |
| 119 | 115        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-115-(10/11)-P     | BA-01-00458 |   | 89      |
| 120 | 119        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-119-(10/11)-P     | BA-01-00449 |   | 140     |
| 121 | 151        | 3     | 50     | 10/11 | 30-Jun-01      | BA1-151-(10/11)-P     | BA-01-00484 |   | 220     |
| 122 | 121        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-121-(10/11)-P     | BA-01-00457 |   | 92      |
| 123 | 136        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-136-(10/11)-P     | BA-01-00453 |   | 160     |
| 124 | 140        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-140-(10/11)-P     | BA-01-00460 |   | 330     |
| 125 | 148        | 4     | 75     | 10/11 | 30-Jun-01      | BA1-148-(10/11)-P     | BA-01-00477 |   | 360     |
| 126 | 104        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-104-(10/11)-P     | BA-01-00450 |   | 650     |
| 127 | 106        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-106-(10/11)-P     | BA-01-00467 |   | 640     |
| 128 | 128        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-128-(10/11)-P     | BA-01-00476 |   | 200     |
| 129 | 155        | 5     | 25     | 10/11 | 30-Jun-01      | BA1-155-(10/11)-P     | BA-01-00483 |   | 880     |
| 130 | 103        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-103-(10/11)-P     | BA-01-00485 |   | 1500    |
| 131 | 110        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-110-(10/11)-P     | BA-01-00482 |   | 1300    |
| 132 | 116        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-116-(10/11)-P     | BA-01-00468 |   | 2400    |
| 133 | 142        | 6     | 50     | 10/11 | 30-Jun-01      | BA1-142-(10/11)-P     | BA-01-00466 |   | 1900    |
| 134 | 120        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-120-(10/11)-P     | BA-01-00474 |   | 2700    |
| 135 | 125        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-125-(10/11)-P     | BA-01-00481 |   | 2400    |
| 136 | 138        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-138-(10/11)-P     | BA-01-00480 |   | 5200    |
| 137 | 150        | 7     | 75     | 10/11 | 30-Jun-01      | BA1-150-(10/11)-P     | BA-01-00489 |   | 1700    |
| 138 | 102        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-102-(10/11)-P     | BA-01-00473 |   | 930     |
| 139 | 114        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-114-(10/11)-P     | BA-01-00488 |   | 1000    |
| 140 | 117        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-117-(10/11)-P     | BA-01-00465 |   | 1400    |
| 141 | 126        | 8     | 25     | 10/11 | 30-Jun-01      | BA1-126-(10/11)-P     | BA-01-00469 |   | 1200    |
| 142 | 112        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-112-(10/11)-P     | BA-01-00472 |   | 1600    |
| 143 | 113        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-113-(10/11)-P     | BA-01-00471 |   | 1600    |
| 144 | 135        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-135-(10/11)-P     | BA-01-00470 |   | 1400    |
| 145 | 154        | 9     | 50     | 10/11 | 30-Jun-01      | BA1-154-(10/11)-P     | BA-01-00487 |   | 4200    |
| 146 | 124        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-124-(10/11)-P     | BA-01-00478 |   | 2800    |
| 147 | 133        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-133-(10/11)-P     | BA-01-00447 |   | 2500    |
| 148 | 158        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-158-(10/11)-P     | BA-01-00464 |   | 2300    |
| 149 | 160        | 10    | 75     | 10/11 | 30-Jun-01      | BA1-160-(10/11)-P     | BA-01-00455 |   | 2700    |
| 150 | 2101       | 3     | 50     | 10/11 | 30-Jun-01      | BA1-2101-(10/11)-P    | BA-01-00486 |   | 92      |
| 151 | 2114       | 8     | 25     | 10/11 | 30-Jun-01      | BA1-2114-(10/11)-P    | BA-01-00462 |   | 1000    |
| 152 | 2125       | 7     | 75     | 10/11 | 30-Jun-01      | BA1-2125-(10/11)-P    | BA-01-00459 |   | 2300    |
| 153 | FAsCtrl    |       |        | 10/11 | 30-Jun-01      | BA1-FAsCtrl-(10/11)-P | BA-01-00475 |   | 40      |

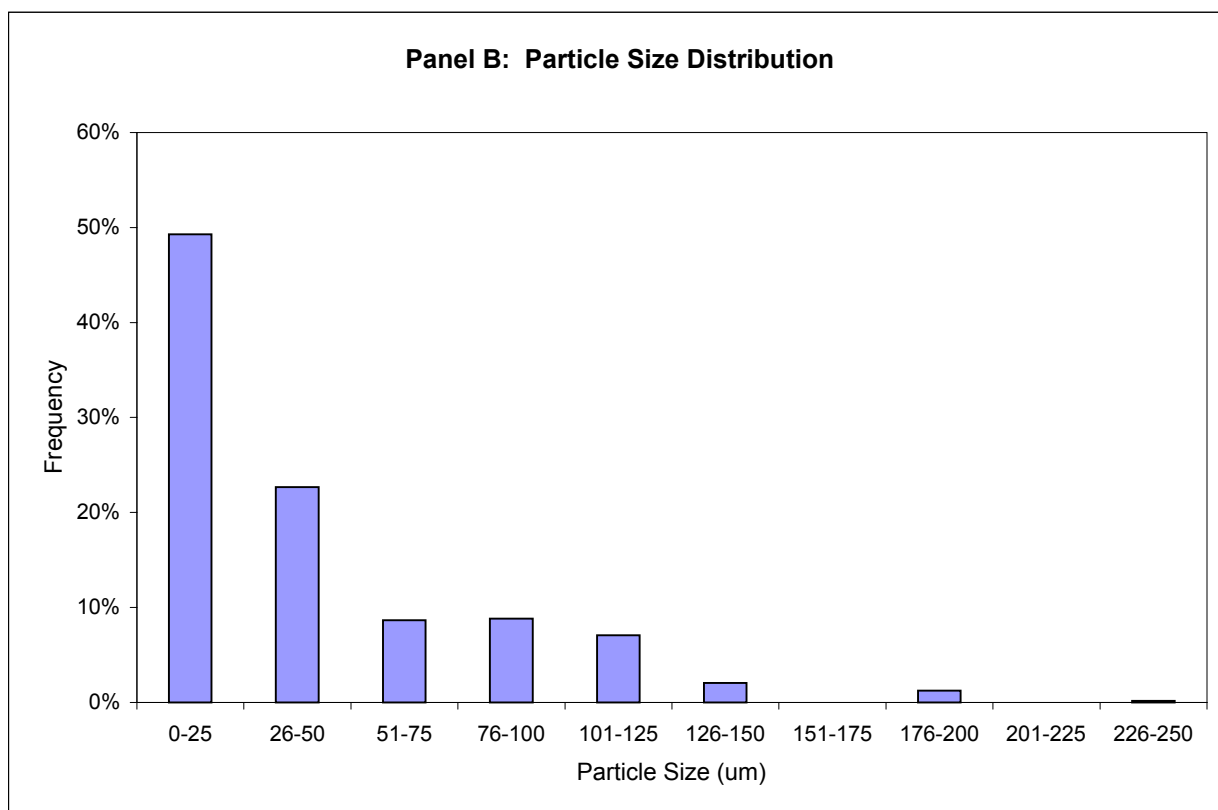
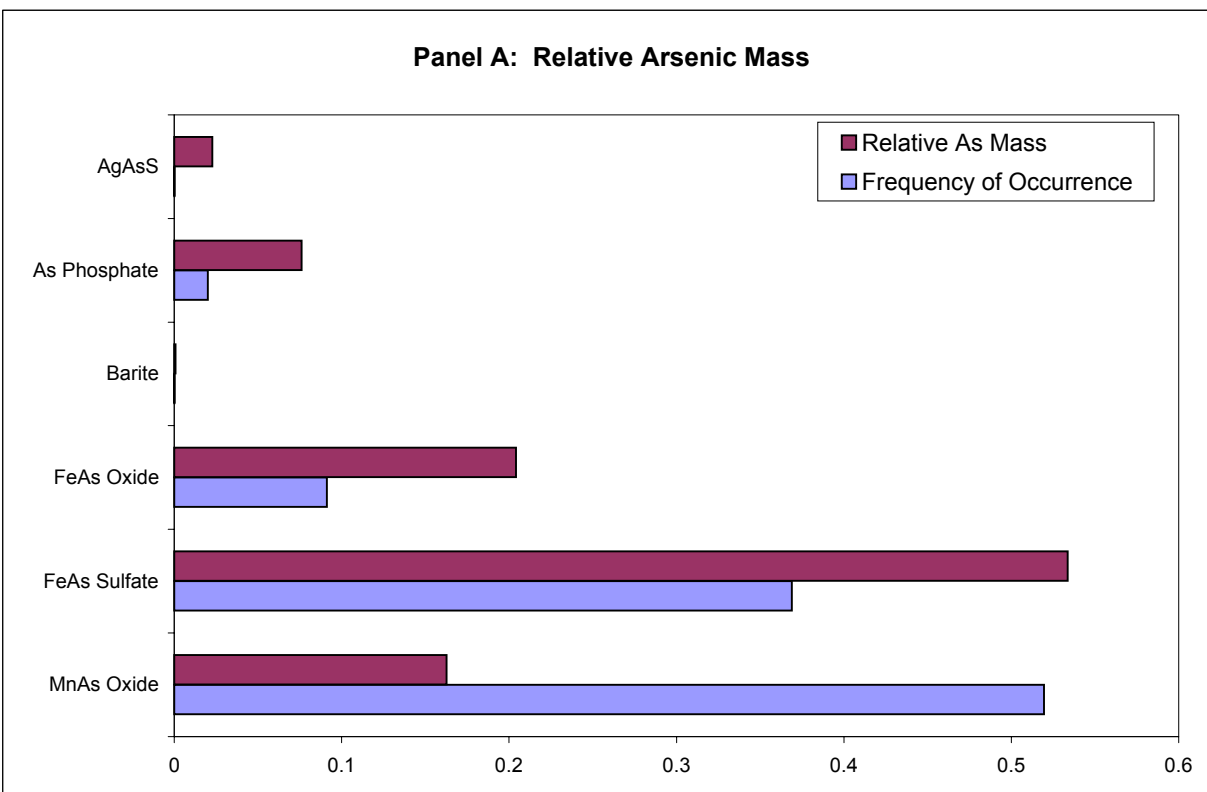
## **APPENDIX B**

### **DETAILED ARSENIC SPECIATION RESULTS**

### Butte Test Material 1 - Arsenic Speciation Summary Statistics

| Phase        | Count |     | Count Frequency (%) |        | Size  |       |      | Size Frequency (%) |        | Relative Arsenic Mass (%) |           |
|--------------|-------|-----|---------------------|--------|-------|-------|------|--------------------|--------|---------------------------|-----------|
|              | Total | Lib | Total               | Lib    | Total | Lib   | Mean | Total              | Lib    | Total                     | Liberated |
| AgAsS        | 1     | 1   | 0.2%                | 0.2%   | 7     | 7     | 7    | 0.02%              | 0.02%  | 2.27%                     | 2.27%     |
| FeAs Oxide   | 22    | 22  | 3.5%                | 3.8%   | 775   | 775   | 35   | 2.34%              | 2.34%  | 9.95%                     | 9.95%     |
| FeAs Oxide   | 37    | 27  | 5.8%                | 4.6%   | 2242  | 1770  | 61   | 6.77%              | 5.35%  | 10.47%                    | 8.26%     |
| MnAs Oxide   | 79    | 79  | 12.4%               | 13.5%  | 10193 | 10193 | 129  | 30.78%             | 30.78% | 11.15%                    | 11.15%    |
| MnAs Oxide   | 161   | 150 | 25.3%               | 25.6%  | 7009  | 6264  | 44   | 21.17%             | 18.92% | 5.11%                     | 4.57%     |
| As Phosphate | 4     | 4   | 0.6%                | 0.7%   | 18    | 18    | 5    | 0.05%              | 0.05%  | 0.05%                     | 0.05%     |
| As Phosphate | 12    | 1   | 1.9%                | 0.2%   | 647   | 12    | 54   | 1.95%              | 0.04%  | 7.55%                     | 0.14%     |
| FeAs Sulfate | 74    | 74  | 11.6%               | 12.6%  | 2889  | 2889  | 39   | 8.72%              | 8.72%  | 15.60%                    | 15.60%    |
| FeAs Sulfate | 245   | 226 | 38.5%               | 38.6%  | 9327  | 8673  | 38   | 28.17%             | 26.19% | 37.76%                    | 35.12%    |
| Barite       | 1     | 1   | 0.2%                | 0.2%   | 5     | 5     | 5    | 0.02%              | 0.02%  | 0.07%                     | 0.07%     |
|              | 636   | 585 | 100.0%              | 100.0% | 33112 | 30606 |      | 100.00%            | 92.43% | 100.00%                   | 87.19%    |

## BUTTE TEST MATERIAL 1 - SPECIATION AND PARTICLE SIZE DATA



### Butte Test Material 2 - Arsenic Speciation Summary Statistics

| Phase        | Count |     | Count Frequency (%) |        | Size  |      |      | Size Frequency (%) |         | Relative Arsenic Mass (%) |           |
|--------------|-------|-----|---------------------|--------|-------|------|------|--------------------|---------|---------------------------|-----------|
|              | Total | Lib | Total               | Lib    | Total | Lib  | Mean | Total              | Lib     | Total                     | Liberated |
| Clays        | 1     | 1   | 0.7%                | 0.7%   | 90    | 90   | 90   | 2.01%              | 2.01%   | 0.07%                     | 0.07%     |
| AsMSO4       | 1     | 1   | 0.7%                | 0.7%   | 8     | 8    | 8    | 0.18%              | 0.18%   | 0.33%                     | 0.33%     |
| FeAs Oxide   | 75    | 75  | 54.7%               | 54.7%  | 2770  | 2770 | 37   | 61.97%             | 61.97%  | 39.28%                    | 39.28%    |
| Pyrite       | 7     | 7   | 5.1%                | 5.1%   | 139   | 139  | 20   | 3.11%              | 3.11%   | 0.13%                     | 0.13%     |
| Slag         | 1     | 1   | 0.7%                | 0.7%   | 85    | 85   | 85   | 1.90%              | 1.90%   | 0.04%                     | 0.04%     |
| Sulfosalts   | 20    | 20  | 14.6%               | 14.6%  | 150   | 150  | 8    | 3.36%              | 3.36%   | 41.87%                    | 41.87%    |
| FeAs Sulfate | 32    | 32  | 23.4%               | 23.4%  | 1228  | 1228 | 38   | 27.47%             | 27.47%  | 18.28%                    | 18.28%    |
|              | 137   | 137 | 100.0%              | 100.0% | 4470  | 4470 |      | 100.00%            | 100.00% | 100.00%                   | 100.00%   |

## BUTTE TEST MATERIAL 2 - SPECIATION AND PARTICLE SIZE DATA

